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FILE 'HOME' ENTERED AT 15:52:10 ON 28 APR 2003

=> file dgene

COST IN U.S. DOLLARS

SINCE FILE TOTAL

ENTRY 0.21 SESSION 0.21

FULL ESTIMATED COST

FILE 'DGENE' ENTERED AT 15:52:35 ON 28 APR 2003

FILE LAST UPDATED: 25 APR 2003

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<20030425/UP>

DGENE CURRENTLY CONTAINS 3,513,779 BIOSEQUENCES

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http://www.stn-international.de/training center/bioseq/dgene wm.pdf

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http://www.stn-international.de/training\_center/bioseq/dgene\_help.pdf

>>> DOWNLOAD DGENE BLAST/GETSIM FREQUENTLY ASKED QUESTIONS: http://www.stn-international.de/service/faq/dgenefaq.pdf <<<

=> s ostogenic protein and nonarticular cartilage repair

0 OSTOGENIC

1401487 PROTEIN

0 OSTOGENIC PROTEIN

(OSTOGENIC (W) PROTEIN)

7 NONARTICULAR

22747 CARTILAGE

51008 REPAIR

0 NONARTICULAR CARTILAGE REPAIR

(NONARTICULAR (W) CARTILAGE (W) REPAIR)

L1 0 OSTOGENIC PROTEIN AND NONARTICULAR CARTILAGE REPAIR

=> s osteogenic protein

2004 OSTEOGENIC

1401487 PROTEIN

L2 1127 OSTEOGENIC PROTEIN

(OSTEOGENIC (W) PROTEIN)

=> s nonarticular cartilage

7 NONARTICULAR

22747 CARTILAGE

L3 7 NONARTICULAR CARTILAGE

(NONARTICULAR (W) CARTILAGE)

1411 LARYNX L47 L3 AND LARYNX => s 13 and 14 L5 7 L3 AND L4 => s 15 and 12 7 L5 AND L2 L<sub>6</sub> => s 16 and repair 51008 REPAIR 0 L6 AND REPAIR => d l6 ti abs ibib tot ANSWER 1 OF 7 DGENE (C) 2003 THOMSON DERWENT L6 ΤI Novel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier ANAAY92442 Protein DGENE AB The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci. ACCESSION NUMBER: AAY92442 Protein DGENE TITLE: Novel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier INVENTOR: Vukicevic S; Katic V; Sampath K T PATENT ASSIGNEE: (STYC) STRYKER CORP. PATENT INFO: WO 2000020021 A1 20000413 65p APPLICATION INFO: WO 1999-US17222 19990730 US 1998-103161 PRIORITY INFO: 19981006 DOCUMENT TYPE: Patent LANGUAGE: English OTHER SOURCE: 2000-317644 [27] CROSS REFERENCES: N-PSDB: AAA09361 DESCRIPTION: Human osteogenic protein 1 (OP-1). 1.6 ANSWER 2 OF 7 DGENE (C) 2003 THOMSON DERWENT TT Novel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier ΑN AAY92441 protein DGENE AΒ Generic Sequence 10 contains generic sequence 9 and an N-terminal extension. Generic sequence 9 is a composite amino acid sequence of the following proteins: human OP-1 to -3, human BMP-2 to -6, -9 to -11, Drosophila 60A, Xenopus Vg-1, sea urchin UNIVIN, human CDMP-1 to -3, human and mouse GDF-1, chicken DORSALIN, DPP, Drosophila Screw, mouse NODAL, mouse GDF-8 to -11, human GDF-8, -11, human BMP-15 and rat BMP3b. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote

chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including

tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear,

65p

nose, ribs, invertebral discs, and interarticular menisci. ACCESSION NUMBER: AAY92441 protein DGENE

TITLE:

Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable carrier Vukicevic S; Katic V; Sampath K T

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161

DOCUMENT TYPE: Patent LANGUAGE: English

**INVENTOR:** 

OTHER SOURCE: 2000-317644 [27]

Generic sequence 10, derived from osteogenic DESCRIPTION:

protein family members.

L6 ANSWER 3 OF 7 DGENE (C) 2003 THOMSON DERWENT

Novel methods for repairing a defect in mammalian nonarticular TI cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AAY92440 protein DGENE ΑN

Generic Sequence 9 is a composite amino acid sequence of the following AΒ proteins: human OP-1 to -3, human BMP-2 to -6, -9 to -11, Drosophila 60A, Xenopus Vg-1, sea urchin UNIVIN, human CDMP-1 to -3, human and mouse GDF-1, chicken DORSALIN, DPP, Drosophila Screw, mouse NODAL, mouse GDF-8 to -11, human GDF-8, -11, human BMP-15 and rat BMP3b. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92440 protein **DGENE** 

TITLE: Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable carrier Vukicevic S; Katic V; Sampath K T

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413 65p

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent LANGUAGE: English

INVENTOR:

OTHER SOURCE: 2000-317644 [27]

DESCRIPTION: Generic sequence 9, derived from osteogenic

protein family members.

L6 ANSWER 4 OF 7 DGENE (C) 2003 THOMSON DERWENT

Novel methods for repairing a defect in mammalian nonarticular TIcartilage tissue or ligaments using an osteogenic

protein in a biocompatible, bioresorbable carrier

AN AAY92439 protein DGENE

AB Generic Sequence 8 contains generic sequence 7 (AAY92438), which accomodates the homologies shared among osteogenic protein family members, including OP-1, OP-2, OP-3, BMP-2 to -6, 60A, DPP, Vg-1, Vgr-1 and GDF, as well as an N-terminal addition of 5 residues. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92439 protein DGENE

Novel methods for repairing a defect in mammalian TITLE:

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable carrier Vukicevic S; Katic V; Sampath K T

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413 65p

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006
DOCUMENT TYPE: Patent
LANGUAGE: English

INVENTOR:

INVENTOR:

OTHER SOURCE: 2000-317644 [27]

DESCRIPTION: Generic sequence 8, derived from osteogenic

protein family members.

L6 ANSWER 5 OF 7 DGENE (C) 2003 THOMSON DERWENT

TINovel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AAY92438 protein AN **DGENE** 

AB Generic Sequence 7 accomodates the homologies shared among osteogenic protein family members, including OP-1, OP-2, OP-3, BMP-2 to -6, 60A, DPP, Vg-1, Vgr-1 and GDF. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92438 protein DGENE

TITLE: Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable carrier Vukicevic S; Katic V; Sampath K T

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413 APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006 65p

DOCUMENT TYPE:

Patent

LANGUAGE:

English

OTHER SOURCE:

2000-317644 [27]

DESCRIPTION:

Generic sequence 7, derived from osteogenic

protein family members.

ANSWER 6 OF 7 DGENE (C) 2003 THOMSON DERWENT L6

Novel methods for repairing a defect in mammalian nonarticular TI

cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AN AAY92437 protein DGENE

AB OPX defines the seven-cysteine skeleton of several OP-1 and OP-2 variants. Each Xaa is chosen from the residues occuring at the corresponding position in the C-terminal sequence of mouse or human OP-1

or OP-2. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which

comprises providing an osteogenic protein in a

biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea

(including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92437 protein DGENE

TITLE:

Novel methods for repairing a defect in mammalian

65p

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable carrier Vukicevic S; Katic V; Sampath K T

INVENTOR:

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO:

WO 2000020021 A1 20000413

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent

English LANGUAGE:

OTHER SOURCE: 2000-317644 [27]

DESCRIPTION: Generic OPX, seven-cysteine skeleton.

L6 ANSWER 7 OF 7 DGENE (C) 2003 THOMSON DERWENT

Novel methods for repairing a defect in mammalian nonarticular ΤI cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AN AAA09361 cDNA **DGENE** 

AB The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including

tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAA09361 cDNA DGENE

TITLE: Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable carrier Vukicevic S; Katic V; Sampath K T

INVENTOR:

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent LANGUAGE: English

LANGUAGE: English
OTHER SOURCE: 2000-317644 [27]
CROSS REFERENCES: P-PSDB: AAY92442

DESCRIPTION: Human osteogenic protein 1 (OP-1) coding

sequence.

#### => d his

(FILE 'HOME' ENTERED AT 15:52:10 ON 28 APR 2003)

FILE 'DGENE' ENTERED AT 15:52:35 ON 28 APR 2003

L1 0 S OSTOGENIC PROTEIN AND NONARTICULAR CARTILAGE REPAIR

65p

L2 1127 S OSTEOGENIC PROTEIN

L3 7 S NONARTICULAR CARTILAGE

L4 7 S L3 AND LARYNX

L5 7 S L3 AND L4 L6 7 S L5 AND L2

L7 0 S L6 AND REPAIR

=> file wpids

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
40.06
40.27

FILE 'WPIDS' ENTERED AT 15:57:11 ON 28 APR 2003 COPYRIGHT (C) 2003 THOMSON DERWENT

FILE LAST UPDATED: 28 APR 2003 <20030428/UP>
MOST RECENT DERWENT UPDATE: 200326 <200326/DW>

DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

Due to data production problems in updates 24 and 25 the WPI file had to be reset to update 200323 on April 24 and the corrected updates were reloaded.

SDIs for update 24 were rerun. The previous SDI run for 24 has been credited.

We also recommend to recreate answer sets dated between April 10 and 24. Charges incurred to accomplish this will be credited of course.

- >>> NEW WEEKLY SDI FREQUENCY AVAILABLE --> see NEWS <<<
- >>> PATENT IMAGES AVAILABLE FOR PRINT AND DISPLAY <<<
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  SEE http://www.derwent.com/dwpi/updates/dwpicov/index.html <<<
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GUIDES, PLEASE VISIT:
http://www.derwent.com/userguides/dwpi guide.html <<<</pre>

=> s l1

0 OSTOGENIC PROTEIN (OSTOGENIC (W) PROTEIN) 3 NONARTICULAR 2940 CARTILAGE 44338 REPAIR O NONARTICULAR CARTILAGE REPAIR (NONARTICULAR (W) CARTILAGE (W) REPAIR) L8O OSTOGENIC PROTEIN AND NONARTICULAR CARTILAGE REPAIR => d his (FILE 'HOME' ENTERED AT 15:52:10 ON 28 APR 2003) FILE 'DGENE' ENTERED AT 15:52:35 ON 28 APR 2003 O S OSTOGENIC PROTEIN AND NONARTICULAR CARTILAGE REPAIR L1 L21127 S OSTEOGENIC PROTEIN L37 S NONARTICULAR CARTILAGE T.4 7 S L3 AND LARYNX L5 7 S L3 AND L4 7 S L5 AND L2 Lб L7 0 S L6 AND REPAIR FILE 'WPIDS' ENTERED AT 15:57:11 ON 28 APR 2003 L8 0 S L1 => s 12 487 OSTEOGENIC 96319 PROTEIN Ь9 91 OSTEOGENIC PROTEIN (OSTEOGENIC (W) PROTEIN) => s 133 NONARTICULAR 2940 CARTILAGE L10 1 NONARTICULAR CARTILAGE (NONARTICULAR (W) CARTILAGE) => s 14 3 NONARTICULAR 2940 CARTILAGE 1 NONARTICULAR CARTILAGE (NONARTICULAR (W) CARTILAGE) 901 LARYNX L111 L3 AND LARYNX => s 153 NONARTICULAR 2940 CARTILAGE 1 NONARTICULAR CARTILAGE (NONARTICULAR (W) CARTILAGE) 3 NONARTICULAR 2940 CARTILAGE 1 NONARTICULAR CARTILAGE (NONARTICULAR (W) CARTILAGE) 901 LARYNX L12 1 L3 AND L4 => s 16 3 NONARTICULAR 2940 CARTILAGE 1 NONARTICULAR CARTILAGE (NONARTICULAR (W) CARTILAGE) 3 NONARTICULAR 2940 CARTILAGE

1 NONARTICULAR CARTILAGE (NONARTICULAR (W) CARTILAGE)

901 LARYNX

487 OSTEOGENIC

96319 PROTEIN

91 OSTEOGENIC PROTEIN

(OSTEOGENIC (W) PROTEIN)

1 L5 AND L2 L13

=> d 113 ti abs ibib tot

ANSWER 1 OF 1 WPIDS (C) 2003 THOMSON DERWENT

TINovel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier.

AN 2000-317644 [27] WPIDS

2000-317706 [27] CR

WO 200020021 A UPAB: 20020910

NOVELTY - Repairing a defect in a nonarticular cartilage tissue or a ligament of a mammal, comprising providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus, inducing the formation of functional replacement cartilage, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) an implantable device for repairing a defect in a nonarticular cartilage tissue comprising an osteogenic protein disposed in a devitalized cartilage, a collagen carrier, or a carboxymethylcellulose carrier; and

(2) promoting chondrogenesis at a defect locus in a mammal comprising providing an osteogenic protein in a devitalized cartilage carrier that is configured to fit into the defect locus.

ACTIVITY - Osteogenic; chondrogenic.

MECHANISM OF ACTION - Osteopathic stimulating implant; transplantation.

USE - The methods and implants are useful for repairing or correcting a defect in a nonarticular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, edema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

Dwq.0/0

ACCESSION NUMBER: 2000-317644 [27] WPIDS

CROSS REFERENCE: 2000-317706 [27] DOC. NO. CPI:

C2000-096081

TITLE:

Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier.

DERWENT CLASS: A96 B04 D22

INVENTOR(S): KATIC, V; SAMPATH, K T; VUKICEVIC, S

PATENT ASSIGNEE(S): (STYC) STRYKER CORP; (CREA-N) CREATIVE BIOMOLECULES INC

COUNTRY COUNT: 23

PATENT INFORMATION:

PATENT NO KIND DATE WEEK PG -------

WO 2000020021 A1 20000413 (200027) \* EN

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: AU CA JP US

AU 9952417 A 20000426 (200036)

A1 20010725 (200143) EP 1117422

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE US 2001024823 A1 20010927 (200159)
JP 2002526167 W 20020820 (200258) 70

#### APPLICATION DETAILS:

PATENT NO K	IND	APPLICATION	DATE
WO 2000020021 AU 9952417	A1 A	WO 1999-US17222	19990730
EP 1117422	A1	AU 1999-52417 EP 1999-937624	19990730 19990730
US 2001024823	Al Provisional	WO 1999-US17222 US 1998-103161P	19990730 19981006
	Cont of	WO 1999-US17222 US 2001-828607	19990730 20010406
JP 2002526167	W	WO 1999-US17222 JP 2000-573380	19990730 19990730

#### FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9952417 EP 1117422	A Based on Al Based on	WO 200020021 WO 200020021
	67 W Based on	WO 200020021

PRIORITY APPLN. INFO: US 1998-103161P 19981006; US 2001-828607 20010406

#### => d his

(FILE 'HOME' ENTERED AT 15:52:10 ON 28 APR 2003)

	FILE	'DGENE'	ENTERED AT 15:52:35 ON 28 APR 2003
L1		0 S	OSTOGENIC PROTEIN AND NONARTICULAR CARTILAGE REPAIR
L2		1127 S	OSTEOGENIC PROTEIN
L3		7 S	NONARTICULAR CARTILAGE
L4		7 S	L3 AND LARYNX
L5		7 S	L3 AND L4
L6		7 S	L5 AND L2
L7		0 S	L6 AND REPAIR
	FILE	'WPIDS'	ENTERED AT 15:57:11 ON 28 APR 2003
L8		0 S	L1
L9		91 S	L2
L10		1 S	L3
L11		1 S	L4
L12		1 S	L5
L13		1 S	L6
=> f	ile ho	anlus	

=> file hcaplus

COST IN U.S. DOLLARS
SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST
6.64
46.91

FILE 'HCAPLUS' ENTERED AT 15:57:59 ON 28 APR 2003
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FILE COVERS 1907 - 28 Apr 2003 VOL 138 ISS 18 FILE LAST UPDATED: 27 Apr 2003 (20030427/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 16

19 NONARTICULAR

19446 CARTILAGE

5 NONARTICULAR CARTILAGE

(NONARTICULAR (W) CARTILAGE)

19 NONARTICULAR

19446 CARTILAGE

5 NONARTICULAR CARTILAGE

(NONARTICULAR (W) CARTILAGE)

2142 LARYNX

2601 OSTEOGENIC

1503618 PROTEIN

266 OSTEOGENIC PROTEIN

(OSTEOGENIC (W) PROTEIN)

L14

1 L5 AND L2

=> d l14 ti abs ibib tot

L14 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2003 ACS

TI Osteogenic proteins for repair of larynx, trachea, and other fibrocartilaginous tissues

AB Provided herein are methods and devices for inducing the formation of functional replacement nonarticular cartilage tissues and ligament tissues. These methods and devices involve the use of osteogenic proteins, and are useful in repairing defects in the larynx, trachea, interarticular menisci, intervertebral disks, ear, nose, ribs and other fibrocartilaginous tissues in a mammal.

ACCESSION NUMBER:

2000:240976 HCAPLUS

DOCUMENT NUMBER:

132:284278

TITLE:

Osteogenic proteins for repair of larynx,

trachea, and other fibrocartilaginous tissues

INVENTOR(S):

Vukicevic, Slobodan; Katic, Vladimir; Sampath, Kuber

т.

PATENT ASSIGNEE(S):

SOURCE:

Stryker Corporation, USA PCT Int. Appl., 65 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	<del>-</del>			
WO 2000020021	<b>A</b> 1	20000413	WO 1999-US17222	19990730
W: AU, CA,	JP, US			
RW: AT, BE,	CH, CY	, DE, DK, ES,	FI, FR, GB, GR, IE	, IT, LU, MC, NL,
PT, SE				
CA 2343698	AA	20000413	CA 1999-2343698	19990730
AU 9952417	A1	20000426	AU 1999-52417	19990730

A1 20010725 EP 1999-937624 19990730 EP 1117422 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI JP 2002526167 19990730 T2 20020820 JP 2000-573380 US 2002106362 A1 20020808 US 2001024823 A1 20010927 US 1999-366021 19990802 US 2001-828607 20010406 US 1998-103161P P 19981006 PRIORITY APPLN. INFO.: WO 1999-US17222 W 19990730 REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT => d his (FILE 'HOME' ENTERED AT 15:52:10 ON 28 APR 2003) FILE 'DGENE' ENTERED AT 15:52:35 ON 28 APR 2003 L1O S OSTOGENIC PROTEIN AND NONARTICULAR CARTILAGE REPAIR L21127 S OSTEOGENIC PROTEIN L37 S NONARTICULAR CARTILAGE 7 S L3 AND LARYNX L47 S L3 AND L4 L5 7 S L5 AND L2 L6L70 S L6 AND REPAIR FILE 'WPIDS' ENTERED AT 15:57:11 ON 28 APR 2003 0 S L1 L891 S L2 Ь9 1 S L3 L10L11 1 S L4 1 S L5 L12L13 1 S L6 FILE 'HCAPLUS' ENTERED AT 15:57:59 ON 28 APR 2003 1 S L6 L14 => file uspatful SINCE FILE TOTAL ENTRY SESSION COST IN U.S. DOLLARS FULL ESTIMATED COST 4.67 51.58 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) TOTAL SINCE FILE ENTRY SESSION CA SUBSCRIBER PRICE -0.65 -0.65 FILE 'USPATFULL' ENTERED AT 15:58:22 ON 28 APR 2003 CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS) FILE COVERS 1971 TO PATENT PUBLICATION DATE: 24 Apr 2003 (20030424/PD) FILE LAST UPDATED: 24 Apr 2003 (20030424/ED) HIGHEST GRANTED PATENT NUMBER: US6553568 HIGHEST APPLICATION PUBLICATION NUMBER: US2003079264 CA INDEXING IS CURRENT THROUGH 24 Apr 2003 (20030424/UPCA) ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 24 Apr 2003 (20030424/PD) REVISED CLASS FIELDS (/NCL) LAST RELOADED: Feb 2003 USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Feb 2003 >>> USPAT2 is now available. USPATFULL contains full text of the <<< >>> original, i.e., the earliest published granted patents or <<< >>> applications. USPAT2 contains full text of the latest US <<< >>> publications, starting in 2001, for the inventions covered in >>> USPATFULL. A USPATFULL record contains not only the original <<< <<< >>> published document but also a list of any subsequent <<< >>> publications. The publication number, patent kind code, and

publication date for all the US publications for an invention <<< >>> are displayed in the PI (Patent Information) field of USPATFULL <<< records and may be searched in standard search fields, e.g., /PN, <<< >>> /PK, etc. >>> USPATFULL and USPAT2 can be accessed and searched together <<< through the new cluster USPATALL. Type FILE USPATALL to <<< >>> enter this cluster. <<< >>> <<< >>> Use USPATALL when searching terms such as patent assignees, <<< >>> classifications, or claims, that may potentially change from <<< >>> the earliest to the latest publication. <<< This file contains CAS Registry Numbers for easy and accurate substance identification. => s 11 8 OSTOGENIC 138123 PROTEIN 1 OSTOGENIC PROTEIN (OSTOGENIC (W) PROTEIN) 17 NONARTICULAR 11218 CARTILAGE 109530 REPAIR O NONARTICULAR CARTILAGE REPAIR (NONARTICULAR (W) CARTILAGE (W) REPAIR) L15 O OSTOGENIC PROTEIN AND NONARTICULAR CARTILAGE REPAIR => s 1617 NONARTICULAR 11218 CARTILAGE 6 NONARTICULAR CARTILAGE (NONARTICULAR (W) CARTILAGE) 17 NONARTICULAR 11218 CARTILAGE 6 NONARTICULAR CARTILAGE (NONARTICULAR (W) CARTILAGE) 2788 LARYNX 2523 OSTEOGENIC 138123 PROTEIN 307 OSTEOGENIC PROTEIN (OSTEOGENIC (W) PROTEIN) 1 L5 AND L2 L16 => d l16 ti abs ibib tot L16 ANSWER 1 OF 1 USPATFULL ΤI Repair of larynx, trachea, and other fibrocartilaginous tissues Provided herein are methods and devices for inducing the formation of AB functional replacement nonarticular cartilage tissues and ligament tissues. These methods and devices involve the use of osteogenic proteins, and are useful in repairing defects in the larynx, trachea, interarticular menisci, intervertebral discs, ear, nose, ribs and other fibrocartilaginous tissues in a mammal. CAS INDEXING IS AVAILABLE FOR THIS PATENT. ACCESSION NUMBER: 2001:165613 USPATFULL TITLE: Repair of larynx, trachea, and other

fibrocartilaginous tissues Vukicevic, Slobodan, Zagreb, Croatia INVENTOR(S):

Katic, Vladimir, Zagreb, Croatia

Sampath, Kuber T., Holliston, MA, United States PATENT ASSIGNEE(S): Creative BioMolecules, Inc. (non-U.S. corporation) NUMBER KIND DATE

PATENT INFORMATION:

US 2001024823 A1 20010927 US 2001-828607 A1 20010406 (9)

APPLICATION INFO.:

RELATED APPLN. INFO.:

Continuation of Ser. No. WO 1999-US17222, filed on 30

Jul 1999, UNKNOWN

NUMBER DATE

PRIORITY INFORMATION:

US 1998-103161P 19981006 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

FISH & NEAVE, 1251 AVENUE OF THE AMERICAS, 50TH FLOOR,

NEW YORK, NY, 10020-1105

NUMBER OF CLAIMS:

1

EXEMPLARY CLAIM: LINE COUNT:

1859

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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FULL ESTIMATED COST

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E3	0	>	VUKICEVIO	C,S/AU
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E5	1		VUKICH D	/AU
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E7	1		VUKICH DA	AVID/AU
E8	1		VUKICH DA	AVID J/AU
E9	1		VUKICH H	'AU
E10	3		VUKICH J	
E11	1		VUKICH J	C/AU
E12	3		VUKICH J	J/AU

=> e katic, v/au

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E1	2	KATIC VLADIMIR VVICA KLAPAN/AU
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E3	0>	KATIC, V/AU
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E5	1	KATICH M J/AU
E6	1	KATICH R/AU
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E8		KATICH STEPHANIE C/AU
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E11	1	KATICHEV A N/AU
E12	1	KATICHEV D I/AU

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L1 2 "KATIC VLADIMIR VVICA KLAPAN"/AU

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E3	0>	SAMPATH, K/AU
E4	1	SAMPATHACHAR K R/AU
E5	1	SAMPATHANUKUL PICHET/AU
E6	1	SAMPATHI L/AU
E7	1	SAMPATHKUM K/AU
E8	1	SAMPATHKUM L/AU
E9	6	SAMPATHKUM P/AU
E10	3	SAMPATHKUM P S/AU
E11	11	SAMPATHKUMAR A/AU
E12	2	SAMPATHKUMAR B/AU

#### => d l1 ti abs ibib tot

L1 ANSWER 1 OF 2 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

TI Acute upper respiratory tract infections and indications for tonsillectomy in children. I. Immunoglobulin synthesis in the palatine tonsil tissue.

AB Current viewpoints and practice concerning indications for tonsillectomy are presented. The annual specific risk for upper respiratory infection in children aged up to 15 is 1.1. The risk is higher in the youngest age group, in whom it rises to 1.8, decreasing with age and being lowest among children aged 12-15 years (0.5). The proportion of tonsillitis among acute upper respiratory tract infections is highest in the age group up to 3 years (36.9%); at the age of 4-5 years it is 37.1%, and is lowest among children aged 12-15 years (21.9%). The risk of tonsillitis caused by streptococci is highest among children aged up to 5 years. Statistical significance of differences in the synthesis of immunoglobulins (G, M, A and sA) and lysozymes in the palatine tonsil tissue of tonsillectomized children and healthy volunteers was tested by non-parametric tests for independent samples. Significant differences of the above mentioned syntheses were found in all entities studied. Any contribution to the documentation on the nature and cause of each tonsillitis in childhood is of great clinical value, because it is the only basis for rational consideration of indications for tonsillectomy.

ACCESSION NUMBER: 1994:315448 BIOSIS DOCUMENT NUMBER: PREV199497328448

TITLE: Acute upper respiratory tract infections and indications

for tonsillectomy in children. I. Immunoglobulin synthesis

in the palatine tonsil tissue.

AUTHOR(S): Katic, Vladimir Vvica Klapan (1); Katic, Milica;

Cvoriscec, Dubravka; Risavi, Ranko; Mercep, Filip Vv Culoerta; Fumic, Ksenija; Fumic, Lidija; Gortan, Damir (1) ENT Dep., Zagreb Univ. Sch. Med., Salata 4, 41000

Zagreb Croatia

SOURCE: International Journal of Pediatric Otorhinolaryngology,

(1994) Vol. 29, No. 3, pp. 169-178.

ISSN: 0165-5876.

DOCUMENT TYPE: Article LANGUAGE: English

CORPORATE SOURCE:

L1 ANSWER 2 OF 2 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC.

TI The role of prostaglandins in growth of squamous head and neck cancer.

In this study we systematically investigated the level of prostaglandin E in blood in correlation with different stages of tumor growth, as well as with the incidence of relapses or metastases in patients suffering from carcinoma of the head and neck. The level of prostaglandin E was assessed by radioimmunoassay test. In the 53 patients with squamous cell carcinoma of the head and neck the average level of prostaglandin E in blood was significantly increased (59.1 +- 32.4 pg/ml) in comparison with the patients not suffering from a malignant disease (34.6 +- 5.4; n = 12) or healthy controls (28.2 +- 4.9; n = 10). The level of prostaglandin E in patients suffering from a malignant disease was found to correlate with the stage of the tumor disease; the percentage of patients in whom the

level of prostaglandin E was higher than the average +2 standard deviation was 87 in stage IV, 47 in stage III and 17 in stage II. Within 15-30 days after tumor removal the level of prostaglandin E generally decreased, but neared the control values only in stages II and III. Furthermore, this level was found to increase during relapse of the disease, mostly in patients with increased preoperative level of prostaglandin E.

ACCESSION NUMBER:
DOCUMENT NUMBER:

1993:96791 BIOSIS PREV199395051987

TITLE:

The role of prostaglandins in growth of squamous head and

neck cancer.

AUTHOR (S):

Katic, Vladimir Vvica Klapan (1); Culo, Filip;

Bukovec, Zeljka; Cuk, Viseslav (1)

CORPORATE SOURCE:

(1) Dep. ENT, Sch. Med., Univ. Zagreb, 41000 Zagreb

SOURCE:

Acta Facultatis Medicae Fluminensis, (1992) Vol. 17, No.

1-2, pp. 7-11.

ISSN: 0065-1206.

DOCUMENT TYPE:

Article

LANGUAGE:

English

SUMMARY LANGUAGE:

English; Serbo-Croatian

# WEST

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## Search Results - Record(s) 1 through 5 of 5 returned.

1. Document ID: US 6475753 B1

L7: Entry 1 of 5

File: USPT

Nov 5, 2002

US-PAT-NO: 6475753

DOCUMENT-IDENTIFIER: US 6475753 B1

TITLE: 94 Human Secreted Proteins

DATE-ISSUED: November 5, 2002

#### INVENTOR-INFORMATION:

NAME	CITY	STATE	ZIP CODE	COUNTRY
Ruben; Steven M.	Olney	MD		
Ni; Jian	Rockville	MD		
Rosen; Craig A.	Laytonsville	MD		
Wei; Ying-Fei	Berkeley	CA		
Young; Paul	Gaithersburg	MD		
Florence; Kimberly	Rockville	MD		
Soppet; Daniel R.	Centreville	VA		
Brewer; Laurie A.	St. Paul	MN		
Endress; Gregory A.	Potomac	MD		
Carter; Kenneth C.	Potomac	MD		
Mucenski; Michael	Cincinnati	OH		
Ebner; Reinhard	Gaithersburg	MD		
Lafleur; David W.	Washington	DC		
Olsen; Henrik	Gaithersburg	MD		
Shi; Yanggu	Gaithersburg	MD		
Moore; Paul A.	Germantown	MD		
Komatsoulis; George	Silver Spring	MD		

US-CL-CURRENT: 435/69.1; 435/252.3, 435/320.1, 435/325, 435/471, 435/69.4, 435/71.1, 530/350, 536/23.5

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KMC
Draw, D	esc Ir	nage									

2. Document ID: US 6258778 B1

L7: Entry 2 of 5

File: USPT

Jul 10, 2001

US-PAT-NO: 6258778

DOCUMENT-IDENTIFIER: US 6258778 B1

TITLE: Methods for accelerating bone and cartilage growth and repair

DATE-ISSUED: July 10, 2001

INVENTOR-INFORMATION:

NAME

CITY

STATE ZIP CODE

COUNTRY

Rodgers; Kathleen E.

Long Beach

CA

DiZerega; Gere S.

Pasedena

CA

US-CL-CURRENT: 514/2; 424/185.1, 514/12, 514/21, 530/300, 530/324

Full Title Citation Front Review Classification Date Reference Sequences Attachments KMC Draw, Desc Image

3. Document ID: US 6171610 B1

L7: Entry 3 of 5

File: USPT

Jan 9, 2001

US-PAT-NO: 6171610

DOCUMENT-IDENTIFIER: US 6171610 B1

\*\* See image for Certificate of Correction \*\*

TITLE: Guided development and support of hydrogel-cell compositions

DATE-ISSUED: January 9, 2001

INVENTOR-INFORMATION:

NAME

CITY

STATE ZIP CODE

COUNTRY

Vacanti; Charles A.

Uxbridge Winchester MA MA

Vacanti; Joseph P. Vacanti; Martin P.

Westborough

MA

US-CL-CURRENT: 424/426

Full Title Citation Front Review Classification Date Reference Sequences Attachments

Draws Desc | Image |

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4. Document ID: US 6027744 A

L7: Entry 4 of 5

File: USPT

Feb 22, 2000

US-PAT-NO: 6027744

DOCUMENT-IDENTIFIER: US 6027744 A

TITLE: Guided development and support of hydrogel-cell compositions

DATE-ISSUED: February 22, 2000

INVENTOR-INFORMATION:

NAME

CITY

STATE ZIP CODE

COUNTRY

Vacanti; Charles A.

Worcester

MA

Vacanti; Joseph P.

Worcester

MA

US-CL-CURRENT: 424/426; 623/13.11, 623/15.12, 623/16.11

ZIP CODE

Full Title Citation Front Review Classification Date Reference Sequences Attachments Draw, Desc Image

KWIC

5. Document ID: US 5736372 A

L7: Entry 5 of 5

File: USPT

Apr 7, 1998

US-PAT-NO: 5736372

DOCUMENT-IDENTIFIER: US 5736372 A

TITLE: Biodegradable synthetic polymeric fibrous matrix containing chondrocyte for in vivo production of a cartilaginous structure

DATE-ISSUED: April 7, 1998

INVENTOR-INFORMATION:

NAME

CITY

STATE

COUNTRY

Vacanti; Joseph P.

Winchester

MA

Vacanti; Charles A.

Lexington

MA

Langer; Robert S.

Newton

MA

US-CL-CURRENT: 435/180; 424/422, 424/426, 424/548, 424/549, 424/93.7, 435/177, 435/178, 435/395, 435/398, 435/402

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	K000
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         Sep 16
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        Oct 24
NEWS 12 Oct 24 Nutraceuticals International (NUTRACEUT) now available on STN
NEWS 13 Nov 18 DKILIT has been renamed APOLLIT
NEWS 14 Nov 25 More calculated properties added to REGISTRY
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                 CSA files on STN
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                 PCTFULL now covers WP/PCT Applications from 1978 to date
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                 TOXCENTER enhanced with additional content
NEWS 18 Dec 17
                 Adis Clinical Trials Insight now available on STN
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                 ENERGY, INSPEC
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                 CANCERLIT is no longer being updated
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        Feb 24 METADEX enhancements
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         Feb 24
                 PCTGEN now available on STN
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                 TEMA now available on STN
         Feb 26 NTIS now allows simultaneous left and right truncation
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NEWS 25 Feb 26 PCTFULL now contains images
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         Mar 24 PATDPAFULL now available on STN
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         Mar 24
                 Additional information for trade-named substances without
                 structures available in REGISTRY
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         Apr 11
                 Display formats in DGENE enhanced
NEWS 32
         Apr 14
                 MEDLINE Reload
NEWS 33
         Apr 17
                 Polymer searching in REGISTRY enhanced
NEWS 34
         Apr 21
                 Indexing from 1947 to 1956 being added to records in CA/CAPLUS
NEWS 35
         Apr 21
                 New current-awareness alert (SDI) frequency in
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=> s osteogenic protein
L1 3240 OSTEOGENIC PROTEIN

=> s BMP or bone morphogenetic protein

L2 25957 BMP OR BONE MORPHOGENETIC PROTEIN

=> s bone morphogenic protein

L3 3010 BONE MORPHOGENIC PROTEIN

=> s 12 and 13

L4 2062 L2 AND L3

=> s 11 and 14

L5 302 L1 AND L4

=> s chondrogenic proteins

L6 14 CHONDROGENIC PROTEINS

=> d 16 ti abs ibib tot

L6 ANSWER 1 OF 14 USPATFULL

TI Compositions for regeneration and repair of cartilage lesions

Disclosed is a cartilage repair product that induces both cell ingrowth into a bioresorbable material and cell differentiation into cartilage tissue. Such a product is useful for regenerating and/or repairing both vascular and avascular cartilage lesions, particularly articular cartilage lesions, and even more particularly mensical tissue lesions, including tears as well as segmental defects. Also disclosed is a method of regenerating and repairing cartilage lesions using such a product.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:26341 USPATFULL

TITLE:

Compositions for regeneration and repair of cartilage

lesions

INVENTOR(S):

Atkinson, Brent, Lakewood, CO, United States

Benedict, James J., Arvada, CO, United States

PATENT ASSIGNEE(S):

Sulzer Biologics, Inc., Austin, TX, United States (U.S.

corporation)

PATENT INFORMATION: APPLICATION INFO.:

US 2000-505209 20000216 (9)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1999-250370, filed

on 16 Feb 1999 Continuation-in-part of Ser. No. WO

1998-EP5100, filed on 12 Aug 1998

DOCUMENT TYPE:

Utility

FILE SEGMENT:

GRANTED

PRIMARY EXAMINER:

Baker, Anne-Marie

LEGAL REPRESENTATIVE:

Sheridan Ross P.C.

NUMBER OF CLAIMS:

41 1

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

14 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT:

3437

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 2 OF 14 USPATFULL

TI Quantitative in vitro bone induction assay

An in vitro assay for quantifying the osteogenic capacity of bone implants involves in vitro isolation and quantitation of specific osteogenic factors. The method disclosed permits direct measurement of the osteogenic capacity of an implant to allow greater predictability of the degree to which new bone will grow in a given area. The method eliminates the need to practice the traditional technique of implanting material into a test animal and subsequently sacrificing the animal to assess bone growth associated with the implant. Since the present method does not involve animal testing, it is an extremely reproducible, rapid, and accurate method for predicting whether an implanted composition or

material will induce bone growth without the need for in vivo assays.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:10633 USPATFULL

TITLE:

Quantitative in vitro bone induction assay Wironen, John F., Alachua, FL, UNITED STATES INVENTOR(S): Jaw, Rebecca, Alachua, FL, UNITED STATES

> KIND DATE NUMBER

PATENT INFORMATION: APPLICATION INFO.:

US 2003008328 A1 20030109 US 2001-897728 A1 20010703 (9)

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE: McAndrews, Held, & Malloy, Ltd., Citicorp Center, 500 West Madison Street, 34th Floor, Chicago, IL, 60661

36

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

1

NUMBER OF DRAWINGS:

19 Drawing Page(s)

LINE COUNT:

1073

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 3 OF 14 USPATFULL 1.6

ΤI Osteogenic devices

AB Disclosed are 1) osteogenic devices comprising a matrix containing osteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; 2) amino acid sequence data, amino acid composition, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, 3) methods of producing osteogenic proteins using recombinant DNA technology, and 4) osteogenically and chondrogenically active synthetic protein constructs.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2001:168097 USPATFULL

TITLE:

Osteogenic devices

INVENTOR (S):

Oppermann, Hermann, Medway, MA, United States Kuberasampath, Thangavel, Medway, MA, United States Rueger, David C., West Roxbury, MA, United States

Ozkaynak, Engin, Milford, MA, United States

PATENT ASSIGNEE(S):

Stryker Corporation, Kalamazoo, MI, United States (U.S.

corporation)

NUMBER KIND DATE -----\_\_\_\_\_

PATENT INFORMATION: APPLICATION INFO.:

US 6297213 B1 20011002 US 1998-74299 19980507 (9)

RELATED APPLN. INFO.:

Continuation of Ser. No. US 1995-417071, filed on 4 Apr 1995, now patented, Pat. No. US 5814604 Continuation of Ser. No. US 1993-145812, filed on 1 Nov 1993, now patented, Pat. No. US 5750651 Division of Ser. No. US 1992-995345, filed on 22 Dec 1992, now patented, Pat. No. US 5258494 Division of Ser. No. US 1989-315342, filed on 23 Feb 1989, now patented, Pat. No. US 5011691 Continuation-in-part of Ser. No. US 1988-232630, filed on 15 Aug 1988, now abandoned Continuation-in-part of Ser. No. US 1988-179406, filed on 8 Apr 1988, now

patented, Pat. No. US 4968590

DOCUMENT TYPE:

Utility

FILE SEGMENT:

GRANTED

PRIMARY EXAMINER:

Kemmerer, Elizabeth

LEGAL REPRESENTATIVE:

Testa, Hurwitz & Thibeault, LLP

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

24

NUMBER OF DRAWINGS:

52 Drawing Figure(s); 30 Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 4 OF 14 USPATFULL L6

Repair of larynx, trachea, and other fibrocartilaginous tissues TI

Provided herein are methods and devices for inducing the formation of AB functional replacement nonarticular cartilage tissues and ligament tissues. These methods and devices involve the use of osteogenic proteins, and are useful in repairing defects in the larynx, trachea, interarticular menisci, intervertebral discs, ear, nose, ribs and other fibrocartilaginous tissues in a mammal.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2001:165613 USPATFULL

TITLE:

Repair of larynx, trachea, and other fibrocartilaginous

INVENTOR(S):

Vukicevic, Slobodan, Zagreb, Croatia Katic, Vladimir, Zagreb, Croatia

Sampath, Kuber T., Holliston, MA, United States

PATENT ASSIGNEE(S):

Creative BioMolecules, Inc. (non-U.S. corporation)

NUMBER KIND DATE \_\_\_\_\_\_ US 2001024823 A1 20010927 US 2001-828607 A1 20010406 (9)

PATENT INFORMATION: APPLICATION INFO.:

RELATED APPLN. INFO.: Continuation of Ser. No. WO 1999-US17222, filed on 30

Jul 1999, UNKNOWN

NUMBER DATE \_\_\_\_\_

PRIORITY INFORMATION:

US 1998-103161P 19981006 (60)

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE: FISH & NEAVE, 1251 AVENUE OF THE AMERICAS, 50TH FLOOR,

NEW YORK, NY, 10020-1105

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

56 1859

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 5 OF 14 USPATFULL 1.6

ΤI Nucleotide sequences encoding osteogenic proteins

Disclosed are 1) osteogenic devices comprising a matrix containing AB osteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; 2) amino acid sequence data, amino acid composition, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, 3) methods of producing osteogenic proteins using recombinant DNA technology, and 4) osteogenically and chondrogenically active synthetic protein constructs.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

INVENTOR (S):

2001:112103 USPATFULL

TITLE:

Nucleotide sequences encoding osteogenic proteins Oppermann, Hermann, Medway, MA, United States Kuberasampath, Thangavel, Medway, MA, United States

Rueger, David C., West Roxbury, MA, United States

Ozkaynak, Engin, Milford, MA, United States

PATENT ASSIGNEE(S):

Stryker Corporation, Kalamazoo, MI, United States (U.S.

corporation)

NUMBER KIND DATE 

PATENT INFORMATION: APPLICATION INFO.:

US 6261835 B1 20010717 US 1995-375901 19950120

19950120 (8)

RELATED APPLN. INFO.: Division of Ser. No. US 1993-145812, filed on 1 Nov

1993, now patented, Pat. No. US 5750651 Division of Ser. No. US 1992-995345, filed on 22 Dec 1992, now patented, Pat. No. US 5258494 Division of Ser. No. US 1989-315342, filed on 23 Feb 1989, now patented, Pat. No. US 5011691 Continuation-in-part of Ser. No. US 1988-232630, filed on 15 Aug 1988, now abandoned Continuation-in-part of Ser. No. US 1988-179406, filed

on 8 Apr 1988, now patented, Pat. No. US 4968590

DOCUMENT TYPE: FILE SEGMENT:

Utility GRANTED

PRIMARY EXAMINER:

Kunz, Gary L. Hayes, Robert C.

ASSISTANT EXAMINER: LEGAL REPRESENTATIVE:

Testa, Hurwitz, Thibeault, LLP

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

52 Drawing Figure(s); 30 Drawing Page(s) 2136

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 6 OF 14 USPATFULL L6

TI Hyaluronan based biodegradable scaffolds for tissue repair

A hyaluronic acid derivitized scaffold and method of forming are AB disclosed. The scaffolds are useful for various medical purposes such as tissue repair, tissue reconstruction and wound healing. In order to enhance these processes the scaffolds may be engineered to incorporate biologically active molecules such as BMP.

1

CAS INDEXING IS AVAILABLE FOR THIS PATENT. ACCESSION NUMBER:

1999:96274 USPATFULL

TITLE:

Hyaluronan based biodegradable scaffolds for tissue

repair

INVENTOR(S):

Valentini, Robert F., Cranston, RI, United States

Kim, Hyun D., Providence, RI, United States

PATENT ASSIGNEE(S):

Brown University, Providence, RI, United States (U.S.

corporation)

NUMBER KIND DATE US 5939323 19990817 US 1997-864709 19970528

PATENT INFORMATION: APPLICATION INFO.:

19970528 (8)

NUMBER DATE -----

PRIORITY INFORMATION:

US 1996-18492P 19960528 (60)

DOCUMENT TYPE:

Utility Granted

FILE SEGMENT: PRIMARY EXAMINER: PRIMARY EXAMINER: Witz, Jean C. ASSISTANT EXAMINER: Hanley, Susan

LEGAL REPRESENTATIVE: Wolf, Greenfield & Sacks, P.C.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

2 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 848

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 7 OF 14 USPATFULL

TT Methods for inducing endochondral bone formation comprising

administering CBMP-2A, CBMP-2B, and/or virants thereof

AΒ Disclosed are 1) osteogenic devices comprising a matrix containing osteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; 2) amino acid sequence data, amino acid composition, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, 3) methods of

producing osteogenic proteins using recombinant DNA technology, and 4) osteogenically and chondrogenically active synthetic protein constructs.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

1998:119121 USPATFULL

TITLE:

Methods for inducing endochondral bone formation comprising administering CBMP-2A, CBMP-2B, and/or

INVENTOR(S):

Oppermann, Hermann, Medway, MA, United States Kuberasampath, Thangavel, Medway, MA, United States Rueger, David C., West Roxbury, MA, United States

Ozkaynak, Engin, Milford, MA, United States

PATENT ASSIGNEE(S):

Stryker Corporation, Natick, MA, United States (U.S.

corporation)

NUMBER KIND DATE -----

PATENT INFORMATION: APPLICATION INFO.:

US 5814604 19980929 US 4170717 19950404 (8)

RELATED APPLN. INFO.:

Continuation of Ser. No. 145812, filed on 1 Nov 1993 which is a division of Ser. No. 995345, filed on 22 Dec 1992, now patented, Pat. No. 5258494 which is a division of Ser. No. 315342, filed on 23 Feb 1989, now patented, Pat. No. 5011691 which is a

continuation-in-part of Ser. No. 232630, filed on 15 Aug 1988, now abandoned which is a continuation-in-part

of Ser. No. 179406, filed on 8 Apr 1988, now

patented, Pat. No. 4968590

DOCUMENT TYPE:

Utility Granted

FILE SEGMENT: PRIMARY EXAMINER:

Jagannathan, Vasu S. ASSISTANT EXAMINER: Jagannathan, Vasu S.

Kemmerer, Elizabeth C.

LEGAL REPRESENTATIVE: Testa, Hurwitz & Thibeault, LLP

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

52 Drawing Figure(s); 30 Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 8 OF 14 USPATFULL 1.6

Cartilage and bone-inducing proteins ΤI

Disclosed are 1) osteogenic devices comprising a matrix containing AB osteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; 2) amino acid sequence data, amino acid composition, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, 3) methods of producing osteogenic proteins using recombinant DNA technology, and 4) osteogenically and chondrogenically active synthetic protein constructs.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

1998:51727 USPATFULL

TITLE:

Cartilage and bone-inducing proteins

INVENTOR(S):

Oppermann, Hermann, Medway, MA, United States Kuberasampath, Thangavel, Medway, MA, United States Rueger, David C., West Roxbury, MA, United States

Ozkaynak, Engin, Milford, MA, United States

PATENT ASSIGNEE(S):

Stryker Corporation, Kalamazoo, MI, United States (U.S.

corporation)

NUMBER KIND DATE -----PATENT INFORMATION: US 5750651 19980512 APPLICATION INFO.: US 1993-145812 19931101 (8)

RELATED APPLN. INFO.: Division of Ser. No. US 1992-995345, filed on 22 Dec

1992, now patented, Pat. No. US 5258494 which is a division of Ser. No. US 1989-315342, filed on 23 Feb 1989, now patented, Pat. No. US 5011691 which is a continuation-in-part of Ser. No. US 1988-232630, filed

on 15 Aug 1988, now abandoned which is a

continuation-in-part of Ser. No. US 1988-179406, filed

on 8 Apr 1988, now patented, Pat. No. US 4968590

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted

PRIMARY EXAMINER: Fitzgerald, David L.
ASSISTANT EXAMINER: Kemmerer, Elizabeth C.

LEGAL REPRESENTATIVE: Testa, Hurwitz & Thibeault, LLP

NUMBER OF CLAIMS: 18 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 52 Drawing Figure(s); 30 Drawing Page(s)

LINE COUNT: 2082

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 9 OF 14 USPATFULL

TI Method for recombinant production of osteogenic protein

Disclosed are 1) osteogenic devices comprising a matrix containing osteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; 2) amino acid sequence data, amino acid composition, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, 3) methods of producing osteogenic proteins using recombinant DNA technology, and 4) osteogenically and chondrogenically active synthetic protein constructs.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 97:86449 USPATFULL

TITLE: Method for recombinant production of osteogenic protein

INVENTOR(S): Oppermann, Hermann, Medway, MA, United States

Kuberasampath, Thangavel, Medway, MA, United States Rueger, David C., West Roxbury, MA, United States

Ozkaynak, Engin, Milford, MA, United States

PATENT ASSIGNEE(S): Stryker Corporation, Natick, MA, United States (U.S.

corporation)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1993-145812, filed on 1 Nov

1993 which is a division of Ser. No. US 1992-995345, filed on 22 Dec 1992, now patented, Pat. No. US 5258494 which is a division of Ser. No. US 1989-315342, filed on 23 Feb 1989, now patented, Pat. No. US 5011691 which is a continuation-in-part of Ser. No. US 1988-232630,

filed on 15 Aug 1988, now abandoned which is a

continuation-in-part of Ser. No. US 1988-179406, filed

on 8 Apr 1988, now patented, Pat. No. US 4968590

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Jacobson, Dian C.
ASSISTANT EXAMINER: Kemmerer, Elizabeth C.

LEGAL REPRESENTATIVE: Testa, Hurwitz & Thibeault, LLP

NUMBER OF CLAIMS: 13 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 49 Drawing Figure(s); 30 Drawing Page(s)

LINE COUNT: 1984

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 10 OF 14 USPATFULL

TI Osteogenic proteins

AB Disclosed are 1) osteogenic devices comprising a matrix containing osteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; 2) amino acid sequence data, amino acid compositions, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, 3) methods of producing osteogenic proteins using recombinant DNA technology, and 4) osteogenically and chondrogenically active synthetic protein constructs.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

94:55637 USPATFULL

TITLE:

Osteogenic proteins

INVENTOR(S):

Oppermann, Hermann, Medway, MA, United States

Kuberasampath, Thangavel, Medway, MA, United States Rueger, David C., West Roxbury MA, United States Ozkaynak, Engin, Milford, MA, United States

PATENT ASSIGNEE(S):

Stryker Corporation, Kalamazoo, MI, United States (U.S.

corporation)

NUMBER KIND DATE -----

PATENT INFORMATION:

US 5324819 19940628 US 1992-950229 19920924 (7)

APPLICATION INFO.: RELATED APPLN. INFO.:

Division of Ser. No. US 1990-621988, filed on 4 Dec

1990, now abandoned which is a division of Ser. No. US 1989-315342, filed on 23 Feb 1989, now patented, Pat. No. US 5011691 which is a continuation-in-part of Ser. No. US 1988-232630, filed on 15 Aug 1988, now abandoned

which is a continuation-in-part of Ser. No. US

1988-179406, filed on 8 Apr 1988, now patented, Pat.

No. US 4968590

DOCUMENT TYPE:

Utility Granted

FILE SEGMENT:

Nutter, Nathan M.

PRIMARY EXAMINER:

LEGAL REPRESENTATIVE: Testa, Hurwitz & Thibeault

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 53 Drawing Figure(s); 30 Drawing Page(s)

LINE COUNT:

1864

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 11 OF 14 USPATFULL

ΤI Osteogenic proteins

AB Disclosed are 1) osteogenic devices comprising a matrix containing osteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; 2) amino acid sequence data, amino acid composition, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, 3) methods of producing osteogenic proteins using recombinant DNA technology, and 4) osteogenically and chondrogenically active synthetic protein constructs.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

93:91748 USPATFULL Osteogenic proteins

TITLE: INVENTOR(S):

Oppermann, Hermann, Medway, MA, United States Kuberasampath, Thangavel, Medway, MA, United States

Rueger, David C., West Roxbury, MA, United States Ozkaynak, Engin, Milford, MA, United States

PATENT ASSIGNEE(S):

Stryker Corporation, Kalamazoo, MI, United States (U.S.

corporation)

KIND DATE NUMBER -----US 5258494 PATENT INFORMATION: 19931102 US 1992-995345 APPLICATION INFO.: 19921222 (7) RELATED APPLN. INFO.: Continuation of Ser. No. US 1990-621988, filed on 4 Dec

> 1990, now abandoned which is a division of Ser. No. US 1989-315342, filed on 23 Feb 1989, now patented, Pat. No. US 5011691 which is a continuation-in-part of Ser. No. US 1988-232630, filed on 15 Aug 1988, now abandoned

which is a continuation-in-part of Ser. No. US

1988-179406, filed on 8 Apr 1988, now patented, Pat.

No. US 4968590

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER:

Nutter, Nathan M.

LEGAL REPRESENTATIVE:

Testa, Hurwitz & Thibeault

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

17

NUMBER OF DRAWINGS:

53 Drawing Figure(s); 30 Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 12 OF 14 USPATFULL L6

Osteogenic proteins ΤI

Disclosed are 1) osteogenic devices comprising a matrix containing ABosteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; 2) amino acid sequence data, amino acid composition, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, 3) methods of producing osteogenic proteins using recombinant DNA technology, and 4) osteogenically and chondrogenically active synthetic protein constructs.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

93:7197 USPATFULL Osteogenic proteins

INVENTOR(S):

TITLE:

Oppermann, Hermann, Medway, MA, United States Kuberasampath, Thangavel, Medway, MA, United States Rueger, David C., West Roxbury, MA, United States

Ozkaynak, Engin, Milford, MA, United States

PATENT ASSIGNEE(S):

Stryker Corporation, Kalamazoo, MI, United States (U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION: APPLICATION INFO.:

US 5182365 19930126 US 5182365 19930126 US 1990-621988 19901204 (7)

RELATED APPLN. INFO.:

Division of Ser. No. US 1989-315342, filed on 23 Feb 1989, now patented, Pat. No. US 5011691 which is a

continuation-in-part of Ser. No. US 1988-232630, filed

on 15 Aug 1988, now abandoned which is a

continuation-in-part of Ser. No. US 1988-179406, filed

on 8 Apr 1988, now patented, Pat. No. US 4968590

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER:

Nutter, Nathan M.

LEGAL REPRESENTATIVE:

Testa, Hurwitz & Thibeault

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

16 1

NUMBER OF DRAWINGS:

41 Drawing Figure(s); 30 Drawing Page(s)

LINE COUNT:

1919

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 13 OF 14 USPATFULL L6

TI Osteogenic devices

AB Disclosed are (1) osteogenic devices comprising a matrix containing osteogenic protein and methods of inducing endochondral bone growth in mammals using the devices; (2) amino acid sequence data, amino acid composition, solubility properties, structural features, homologies and various other data characterizing osteogenic proteins, (3) methods of producing osteogenic proteins using recombinant DNA technology, and (4) osteogenically and chondrogenically active synthetic protein constructs.

osteogenically and chondrogenically active synthetic protein constructs. CAS INDEXING IS AVAILABLE FOR THIS PATENT. 91:34212 USPATFULL ACCESSION NUMBER: Osteogenic devices TITLE: INVENTOR(S): Oppermann, Hermann, Medway, MA, United States Kuberasampath, Thangavel, Medway, MA, United States Rueger, David C., West Roxbury, MA, United States Ozkaynak, Engin, Milford, MA, United States Stryker Corporation, Kalamazoo, MI, United States (U.S. PATENT ASSIGNEE(S): corporation) NUMBER KIND DATE US 1989-315342 PATENT INFORMATION: 19910430 APPLICATION INFO.: 19890223 (7) Continuation-in-part of Ser. No. US 1988-232630, filed RELATED APPLN. INFO.: on 15 Aug 1988 which is a continuation-in-part of Ser. No. US 1988-179406, filed on 8 Apr 1988, now patented, Pat. No. US 4968590 DOCUMENT TYPE: Utility FILE SEGMENT: Granted PRIMARY EXAMINER: Nutter, Nathan M. LEGAL REPRESENTATIVE: Lahive & Cockfield NUMBER OF CLAIMS: EXEMPLARY CLAIM: NUMBER OF DRAWINGS: 49 Drawing Figure(s); 28 Drawing Page(s) LINE COUNT: CAS INDEXING IS AVAILABLE FOR THIS PATENT. ANSWER 14 OF 14 WPIDS (C) 2003 THOMSON DERWENT 1.6 ΤI Osteogenic devices comprising matrix contg. osteogenic protein - useful for inducing endochondral bone growth e.g. in none-union fractures. AN 1989-324202 [44] WPIDS CR 1989-324203 [44]; 1990-290311 [38]; 1991-148697 [20]; 1992-167101 [20]; 1992-167153 [20]; 1992-331475 [40]; 1993-100652 [12]; 1993-100993 [12]; 1993-117208 [14]; 1993-395405 [49]; 1994-007210 [01]; 1994-065304 [08]; 1994-065399 [08]; 1994-065689 [08]; 1994-118107 [14]; 1994-118121 [14]; 1994-118146 [14]; 1994-118148 [14]; 1994-167392 [20]; 1994-302971 [37]; 1994-324521 [40]; 1996-010159 [01]; 1997-178399 [16]; 1997-384665 [35]; 1998-109345 [10]; 1998-158353 [14]; 1998-260496 [23]; 1998-333785 [30]; 1999-131303 [11]; 1999-589530 [50]; 2000-038265 [03]; 2000-422077 [36]; 2001-069971 [08]; 2002-415042 [44] AB 8909787 A UPAB: 20030407 Osteogenic device for implantation in a mammal comprises a biocompatible, in vivo biodegradable matrix (I), defining pores of sufficient dimension to permit influx, proliferation and differentiation of migratory progenitor cells from the body, and, disposed in the matrix and accessible to the cells, pure osteogenic protein (II), which is capable of inducing

endochondrial bone formation in the mammal.

Also claimed is a DNA sequence encoding a protein which induces bone or cartilage formation when implanted in a mammal in association with a matrix. The novel DNA is duplicative of a gene of defined sequence.

(I) comprises demineralised, protein-extd., particulate, allogenic bone, or demineralised, protein-extd., particulate, deglycosylated xenogenic bone. In glycosylated form, (II) has an apparent mol. wt. of 30 kD (as determined by SDS-polyacrylamide gel electrophoresis). Redn. yields two polypeptides of mol. wts. 16 and 18 kD. In unglycosylated form, (II) has an apparent mol. wt. of 27 kD, on redn. yielding polypeptides of 14 and 16 kD.

USE/ADVANTAGE - The efficacy of bone-inducing potential of the devices was tested in cat and rabbit models and found to be potent

inducers of osteogenesis, ultimately resulting in formation of mineralised bone. Clinical applicns. include correction of acquired and congenital craniofacial and other skeletal or dental anomalies, induction of local endochondrial formation in non-union fractures, periodontal applicns. requiring bone formation, and cartilage repair, e.g. in the treatment of osteoarthritis. (II) has a half-max. bone forming activity of 0.8-1.0 ng/mg of implant.

Dwg.0/19

ABEO US 4968590 A UPAB: 19930923

Pure mammalian osteogenic protein induces endochondral bone formation in association with a matrix upon implantation. Osteogenic protein has half max. activity of 25-50 ng per 25 mg of matrix.

Protein has apparent mol.wt. of 30kD when oxidised w.r.t. mol.wt. standards in SDS-polyacrylamide gel electrophoresis, and comprises 2 separate polypeptide chains, each of apparent mol.wt. 16kD and 18kD respectively.

USE/ADVANTAGE - Can be rapidly and reproducibly purified from mammalian bone, for bone repair procedures. @@

ABEQ US 5011691 A UPAB: 19930923

Osteogenic implant device comprises a biocompatible polymer matrix which is degradable in vivo and which contains a dispersion of osteogenic protein; such that the matrix pores allow the ingress, proliferation and differentiation of migratory progenitor cells from the bloodstream. The osteogenic protein is obtd. by expression of suitable recombinant DNA in a host cell, and comprises two polypeptide chains each with an aminoacid sequence that duplicates an adequate part of the sequence in COP-5 or COP-7, so that after dimerisation by means of disulphide linkage, the resulting conformation and the osteogenic protein together induce endochondrial bone formation.

USE - The prods. accelerate bone formation and healing.

ABEQ US 5108753 A UPAB: 19930923

Osteogenic implant for mammals comprises a biocompatible porous insoluble matrix (constructed from collagen, hydroxyapatite, tricalcium phosphate, polylactic acids, polyglycolic acids, demineralised and guanidine-free allogenic bond, or their mixts.) which allows the influx, differentiation and proliferation of migratory progenitor cells from the body; on the surface of which osteogenic protein is immobilised, with polypeptide chains bonded through disulphide linkages to form dimers having a conformation that induces endochondral bone formation.

% USE - The prods. are osteogenic implants for orthopaedic replacements and repairs.

ABEQ US 5182365 A UPAB: 19930923

Protein produced by expression of recombinant DNA in a host cell, comprising other contaminants, consists of two polypeptide chains each having less than 200 amino acids. Sequence is sufficiently duplicative of that of COP-5 or COP-7 so that the chains, when S-S bonded, has a conformation capable of inducing bore or cartilage formation in association with a matrix when implanted in a mammal.

USE/ADVANTAGE - Useful in xenogenic implants to induce osteogenesis and to repair bone and cartilage. 0/22

ABEQ US 5250302 A UPAB: 19931130

DNA sequence hybridises with characteristic DNA probe sequence, and encodes a polypeptide chain which when expressed, associates with a second polypeptide chain. Chains oxidise to form a dimeric protein species.

Dimeric protein species has a half max. bone-forming activity of 25-50 mg per 25 mg matrix upon implantation to induce endochondral bone formation in a mammal.

ADVANTAGE - Osteogenic protein is rapidly and reproducibly purified from mammalian bone. Osteogenic device prepd. repairs bone. Dwg.0/14

ABEQ US 5258494 A UPAB: 19931220

Nucleic acid (cDNA) that encodes the prodn. of osteogenic and chondrogenic proteins, and plasmids and expression

vectors contg. this DNA are new. Mammalian host cells have been transformed with these expression vectors and then propogated to produce the exogenous proteins.

The proteins have Mr about 27,000-30,000; comprise a pair of polypeptide chains each contg. up to 200 aminoacid units, linked through a disulphide bridge; and are opt. glycosylated. The nucleotide sequence of the cDNA and the aminoacid sequence of the proteins have been defined.

USE - The prods. are mounted in matrixes for osteogenic implants, to induce endochondral bone and cartilage growth.

Dwg.0/22

ABEQ US 5324819 A UPAB: 19940810

Protein produced by expression of recombinant DNA comprises 2 polypeptide chains -S-S- bonded, where one has less than 200 amino acids and the sequence given in the specification. Protein induces bone and cartilage formation in assoc. with a matrix when implanted into a mammal.

USE/ADVANTAGE - For inducing full developmental cascade of endochandral bone formation and bone marrow development. Dwg.0/3

ABEQ EP 411105 B UPAB: 19950727

A matrix for implantation in a mammalian host comprising biodegradable, biocompatible, mineral-free, delipidated Type I insoluble bone collagen particles xenogenic to said host having intraparticle pores and having a mean particle diameter within the range of 70 micron to 850 micron, said particles being substantially depleted in noncollagenous protein and treated with a collagen fibril modifying agent to increase the surface area, pore number and intrusion volume of said particles.

Dwg.0/5

ABEQ EP 362367 B UPAB: 19960329

Use of a compsn. consisting essentially of a single species of osteogenic protein as active osteogenic ingredient, the protein comprising a pair of polypeptide chains bonded in the unreduced state to form a homo- or heterodimeric species having a conformation such that the pair of polypeptide chains is capable of inducing endochondral bone formation when disposed within a matrix and implanted in a mammal, for the mfr. of a medicament for inducing endochondral bone formation.

Dwg.0/19

ABEQ US 5496552 A UPAB: 19960417

An osteogenic device for implantation in a mammal, the device comprising: a porous biocompatible matrix; and

substantially pure osteogenic protein disposed in said matrix and comprising a pair of polypeptide chains disulphide bonded to produce a dimeric species having a conformation such that said pair of polypeptide chains is capable of inducing endochondral bone formation in a mammal when disposed within said porous matrix and implanted in a mammal. Dwg.0/23

ABEQ US 5670336 A UPAB: 19971105

A method for producing an OP-1 protein comprising the step of transforming a cell with a vector having inserted therein a DNA sequence which encodes an amino acid sequence comprising:

 ${\tt LYVSFRDLGWQDWIIAPEGYAAYYCEGECAFPLNSYMNATNHAIVQTLVHFINPETVPKPCC} \\ {\tt APTQLNAISVLYFDDSSNVILKKYRNMVVRACGCH,} \\$ 

culturing said cells in a suitable culture medium, and isolating said OP-1 protein produced by said cell.

Dwg.0/22

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Dwg.0/22
ACCESSION NUMBER: 1989-324202 [44] WPIDS
CROSS REFERENCE: 1989-324203 [44]; 1990-290311 [38]; 1991-148697 [20]; 1992-167101 [20]; 1992-167153 [20]; 1992-331475 [40]; 1993-100652 [12]; 1993-100993 [12]; 1993-117208 [14]; 1993-395405 [49]; 1994-065689 [08]; 1994-065304 [08]; 1994-065399 [08]; 1994-065689 [08]; 1994-118107 [14]; 1994-118121 [14]; 1994-118146 [14]; 1994-118148 [14]; 1994-167392 [20]; 1994-302971 [37]; 1994-324521 [40]; 1996-010159 [01]; 1997-178399 [16]; 1997-384665 [35]; 1998-109345 [10]; 1998-158353 [14]; 1998-260496 [23];
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1998-333785 [30]; 1999-131303 [11]; 1999-589530 [50]; 2000-038265 [03]; 2000-422077 [36]; 2001-069971 [08]; 2002-415042 [44] DOC. NO. NON-CPI: N1989-246922 DOC. NO. CPI: C1989-143593 TITLE: Osteogenic devices comprising matrix contg. osteogenic protein - useful for inducing endochondral bone growth e.g. in none-union fractures. DERWENT CLASS: A96 B04 B07 D16 D22 P13 P32 P34 KUBERASAMPATH, T; OPPERMANN, H; OZKAYNAK, E; RUEGER, D C; INVENTOR(S): RIDGE, R J; OPPERMAN, H; KUBERASAMP, T PATENT ASSIGNEE(S): (STYC) STRYKER CORP; (CREA-N) CREATIVE BIOMOLECULES INC COUNTRY COUNT: 33 PATENT INFORMATION: PATENT NO KIND DATE WEEK LΑ \_\_\_\_\_ WO 8909787 A 19891019 (198944)\* EN 104 RW: AT BE CH DE FR GB IT LI LU NL OA SE W: AU BB BG BR DK FI HU JP KP KR LK MC MG MW NO RO SD SU US AU 8934449 A 19891103 (199003) AU 8935305 A 19891103 (199003) EP 362367 A 19900411 (199015) R: AT BE CH DE FR GB IT LI LU NL SE US 4968590 A 19901106 (199047) W 19910214 (199113) JP 03500655 A 19910430 (199119) US 5011691 JP 03502579 W 19910613 (199130) JP 03504736 W 19911017 (199148) A 19920428 (199220) US 5108753 26 US 5182365 A 19930126 (199307) 58 US 5250302 A 19931005 (199341) 28 US 5258494 A 19931102 (199345) 56 US 5324819 A 19940628 (199425) 53 EP 411105 B1 19950621 (199529) EN R: AT BE CH DE DK ES FR GB IT LI LU NL SE EP 362367 B1 19960228 (199613) EN R: AT BE CH DE FR GB IT LI LU NL SE US 5496552 A 19960305 (199615) E 19960404 (199619) DE 68925773 EP 714665 A2 19960605 (199627) R: AT BE CH DE FR GB IT LI LU NL SE EP 723013 A2 19960724 (199634) EN R: AT BE CH DE FR GB IT LI LU NL SE JP 2522568 B2 19960807 (199636) 38 JP 08187084 A 19960723 (199639) 38 CA 1338663 C 19961022 (199702) JP 08322570 A 19961210 (199708) 33 JP 08336390 A 19961224 (199710) 41 US 5670336 A 19970923 (199744) 53 EP 714665 A3 19971203 (199817) US 5750651 A 19980512 (199826) US 5814604 A 19980929 (199846) B2 19990310 (199915) JP 2869381 37 JP 2933867 B2 19990816 (199938) 39 US 6261835 B1 20010717 (200142) B1 20011002 (200160) US 6297213 EP 1221484 A2 20020710 (200253) R: AT BE CH DE FR GB IT LI LU NL SE EP 1225225 A2 20020724 (200256) R: AT BE CH DE FR GB IT LI LU NL SE EP 714665 B1 20030122 (200308) EN R: AT BE CH DE FR GB IT LI LU NL SE

DE 68929453 E 20030227 (200323)

### APPLICATION DETAILS:

PATENT NO	KIND		AP	PLICATION	DATE
EP 362367	A		EP	 1989-904986	19890407
US 4968590	A			1988-179406	19880408
JP 03500655	W		JP	1989-504771	19890407
US 5011691	Α		US	1989-315342	19890223
JP 03502579	W		JP	1989-504777	19890407
JP 03504736	W		JP	1990-504059	19900222
US 5108753	Α			1990-579865	19900907
US 5182365	Α	CIP of		1988-179406	19880408
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		Div ex		1989-315342	19890223
110 505000	70	Di	US		19901204
US 5250302	A	Div ex Div ex	US	1988-179406	19880408
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03 3230494	^	CIP of	US		19880815
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			US	1992-995345	19921222
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EP 411105	В1			1990-904002	19900222
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EP 362367	B1			1989-904986	19890407
HC E406EE2	7	Div ex		1989-US1453 1988-179406	19890407 19880408
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		Cont of		1992-827052	19920128
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DE 68925773	E		DE	1989-625773	19890407
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EP 714665	A2	Div ex	EP	1989-904986	19890407
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EP 723013	A2	Div ex		1989-904959	19890407
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JP 2522568	В2			1989-504771	19890407 19890407
JP 08187084	Α	Div ex		1989-US1453 1989-504777	19890407
UP 0010/004	A	DIA 6Y		1995-263371	19890407
CA 1338663	С			1989-596144	19890407
JP 08322570	A	Div ex		1996-3435	19890407
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JP 08336390	Α	Div ex		1989-504771	19890407
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US 5670336	A	CIP of		1988-179406	19880408
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		Div ex		1989-315342	19890223
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PD 714665	ר ע	Div ex		1995-376731	19950120
EP 714665	A3	DIA 6X		1989-904986 1995-201872	19890407 19890407
US 5750651	A	CIP of		1988-179406	19880408
32 3,30031		CIP of		1988-232630	19880815
		- <del>-</del>			

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			Div ex		1992-995345	19921222
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			Cont of		1995-417071	19950404
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			Div ex		1996-200044	19890407
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			Div ex		1995-201872	19890407
				ΕP	2001-201546	19890407
ΕP	714665	В1	Div ex		1989-904986	19890407
					1995-201872	19890407
			Related		2001-201546	19890407
DE	68929453	E			1989-629453	19890407
				EΡ	1995-201872	19890407

# FILING DETAILS:

PATENT NO	KIND	PATENT NO
US 5108753 US 5182365	A Div ex A CIP of	US 4968590 US 4968590
US 5250302	Div ex A Div ex Div ex	US 5011691 US 4968590 US 5108753
US 5258494	A CIP of Div ex	US 4968590 US 5011691
US 5324819	A CIP of Div ex	US 4968590 US 5011691
EP 411105	B1 Based on	WO 9010018
EP 362367 US 5496552	A Div ex	WO 8909787 US 4968590
	Div ex Cont of	US 5108753 US 5250302
DE 68925773	E Based on Based on	EP 362367 WO 8909787
JP 2522568	B2 Previous Pub Based on	
US 5670336	A CIP of Div ex Div ex	US 4968590 US 5011691 US 5258494
EP 714665	A3 Div ex	EP 362367

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US 5011691
                     Div ex
                                    US 5258494
                     Div ex
                                    US 4968590
     US 5814604
                  A CIP of
                                    US 5011691
                     Div ex
                     Div ex
                                    US 5258494
                  B2 Previous Publ. JP 08322570
     JP 2869381
                B2 Previous Publ. JP 08336390
     JP 2933867
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                                    US 4968590
     US 6261835
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                     Div ex
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                                     US 5258494
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                                     US 5750651
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                                     US 5814604
                                     EP 372031
     EP 1221484
                  A2 Div ex
                                    EP 723013
                     Div ex
                                    EP 362367
                  A2 Div ex
     EP 1225225
                                    EP 714665
                     Div ex
                                    EP 1225225
     EP 714665
                  B1 Related to
                     Div ex
                                    EP 362367
     DE 68929453
                  E Based on
                                    EP 714665
PRIORITY APPLN. INFO: US 1989-315342 19890223; US 1988-179406
                     19880408; US 1988-232630 19880815; US
                     1990-579865 19900907; US 1990-621988
                     19901204; US 1992-827052 19920128; US
                                  19921222; US 1992-950229
                     1992-995345
                     19920924; US 1989-422613 19891017; US
                     1993-103604 19930806; US 1994-268252
                     19940629; US 1993-145812 19931101; US
                     1995-376731 19950120; US 1995-417071
                     19950404; US 1995-375901 19950120; US
                     1998-74299
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     BIOBUSINESS, WPIDS, CEABA-VTB, JAPIO, JICST-EPLUS' ENTERED AT 10:27:23 ON
     28 APR 2003
L1
          3240 S OSTEOGENIC PROTEIN
L2
         25957 S BMP OR BONE MORPHOGENETIC PROTEIN
          3010 S BONE MORPHOGENIC PROTEIN
1.3
L4
          2062 S L2 AND L3
L5
           302 S L1 AND L4
            14 S CHONDROGENIC PROTEINS
L6
=> s cartilage replacement
          237 CARTILAGE REPLACEMENT
=> s 17 and 15
L8
            1 L7 AND L5
=> d l8 ti abs ibib tot
L8
    ANSWER 1 OF 1 USPATFULL
TI
      Compositions and methods for the repair and construction of bone and
      other tissue
AB
      The invention relates to novel compositions comprising genetically
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US 4968590

A CIP of

US 5750651

engineered cells and one or more polymers. In an additional aspect, the present invention relates to a method for repairing tissue, for example, cranioskeletal or maxillary bone defects, comprising tranducing the BMP-2 gene into bone marrow stromal cells which are harvested from a subject, combining the genetically engineered cells with at least one polymer, and implanting the combination at the site of the defect. The BMP-2 protein is advantageously produced as long as the tranduced gene stays in the cells.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2002:322028 USPATFULL

TITLE:

Compositions and methods for the repair and

construction of bone and other tissue

INVENTOR(S):

Chang, Chia Ning (Sophia), Taipei, TAIWAN, PROVINCE OF

CHINA

NUMBER KIND DATE \_\_\_\_\_\_\_

PATENT INFORMATION:

US 2002182189 A1 20021205 US 2001-837217 A1 20010419

APPLICATION INFO .: DOCUMENT TYPE:

A1 20010419 (9)

Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE:

Joseph A. Mahoney, MAYER, BROWN & PLATT, P.O. Box 2828,

Chicago, IL, 60690-2828

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

19 Drawing Page(s)

LINE COUNT:

815

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> s BMP-5

L9 851 BMP-5

=> s 19 and 17

L10 2 L9 AND L7

=> d l10 ti abs ibib tot

L10 ANSWER 1 OF 2 USPATFULL

Device and method for regeneration and repair of cartilage lesions ТT Disclosed is a cartilage repair product that induces both cell ingrowth AB into a bioresorbable material and cell differentiation into cartilage tissue. Such a product is useful for regenerating and/or repairing both vascular and avascular cartilage lesions, particularly articular cartilage lesions, and even more particularly mensical tissue lesions, including tears as well as segmental defects. Also disclosed is a method of regenerating and repairing cartilage lesions using such a product.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:33183 USPATFULL

TITLE:

Device and method for regeneration and repair of

cartilage lesions

INVENTOR(S):

Atkinson, Brent, Lakewood, CO, United States

Benedict, James J., Arvada, CO, United States

PATENT ASSIGNEE(S):

Sulzer Biologics Inc., Austin, TX, United States (U.S.

corporation)

NUMBER KIND DATE -----

PATENT INFORMATION: APPLICATION INFO.:

US 6514514 B1 20030204 US 1999-250370 19990216 19990216 (9)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. WO 1998-EP5100, filed

on 12 Aug 1998

NUMBER DATE

PRIORITY INFORMATION: EP 1997-810567 19970814

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Baker, Anne-Marie LEGAL REPRESENTATIVE: Sheridan Ross P.C.

NUMBER OF CLAIMS: 58
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 14 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 2122

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L10 ANSWER 2 OF 2 USPATFULL

TI Compositions for regeneration and repair of cartilage lesions

Disclosed is a cartilage repair product that induces both cell ingrowth into a bioresorbable material and cell differentiation into cartilage tissue. Such a product is useful for regenerating and/or repairing both vascular and avascular cartilage lesions, particularly articular cartilage lesions, and even more particularly mensical tissue lesions, including tears as well as segmental defects. Also disclosed is a method of regenerating and repairing cartilage lesions using such a product.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2003:26341 USPATFULL

TITLE: Compositions for regeneration and repair of cartilage

lesions

INVENTOR(S): Atkinson, Brent, Lakewood, CO, United States

Benedict, James J., Arvada, CO, United States

PATENT ASSIGNEE(S): Sulzer Biologics, Inc., Austin, TX, United States (U.S.

corporation)

US 2000-505209

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-250370, filed

on 16 Feb 1999 Continuation-in-part of Ser. No. WO

20000216 (9)

1998-EP5100, filed on 12 Aug 1998

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Baker, Anne-Marie LEGAL REPRESENTATIVE: Sheridan Ross P.C.

NUMBER OF CLAIMS: 41 EXEMPLARY CLAIM: 1

APPLICATION INFO.:

NUMBER OF DRAWINGS: 14 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 3437

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, SCISEARCH, FSTA, BIOSIS, BIOBUSINESS, WPIDS, CEABA-VTB, JAPIO, JICST-EPLUS' ENTERED AT 10:27:23 ON 28 APR 2003

L1 3240 S OSTEOGENIC PROTEIN

L2 25957 S BMP OR BONE MORPHOGENETIC PROTEIN

L3 3010 S BONE MORPHOGENIC PROTEIN

L4 2062 S L2 AND L3 L5 302 S L1 AND L4

L6 14 S CHONDROGENIC PROTEINS

237 S CARTILAGE REPLACEMENT L7

L8 1 S L7 AND L5 851 S BMP-5 L9 2 S L9 AND L7 L10

=> s bmp-3

11 FILES SEARCHED... L11 762 BMP-3

=> s 111 and 17

L12 2 L11 AND L7

=> d l12 ti abs ibib tot

L12 ANSWER 1 OF 2 USPATFULL

TI Device and method for regeneration and repair of cartilage lesions Disclosed is a cartilage repair product that induces both cell ingrowth AB into a bioresorbable material and cell differentiation into cartilage tissue. Such a product is useful for regenerating and/or repairing both vascular and avascular cartilage lesions, particularly articular cartilage lesions, and even more particularly mensical tissue lesions, including tears as well as segmental defects. Also disclosed is a method of regenerating and repairing cartilage lesions using such a product.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2003:33183 USPATFULL

Device and method for regeneration and repair of TITLE:

cartilage lesions

Atkinson, Brent, Lakewood, CO, United States INVENTOR(S):

Benedict, James J., Arvada, CO, United States

Sulzer Biologics Inc., Austin, TX, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6514514 B1 20030204 APPLICATION INFO.: US 1999-250370 19990216 (9)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. WO 1998-EP5100, filed

on 12 Aug 1998

NUMBER DATE \_\_\_\_\_

PRIORITY INFORMATION: EP 1997-810567 19970814 DOCUMENT TYPE: Utility

FILE SEGMENT: GRANTED
PRIMARY EXAMINER: Baker, Anne-Marie LEGAL REPRESENTATIVE: Sheridan Ross P.C.

NUMBER OF CLAIMS: 58 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 14 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 2122

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 2 OF 2 USPATFULL

TТ Compositions for regeneration and repair of cartilage lesions

Disclosed is a cartilage repair product that induces both cell ingrowth AB into a bioresorbable material and cell differentiation into cartilage tissue. Such a product is useful for regenerating and/or repairing both vascular and avascular cartilage lesions, particularly articular cartilage lesions, and even more particularly mensical tissue lesions, including tears as well as segmental defects. Also disclosed is a method of regenerating and repairing cartilage lesions using such a product.

ACCESSION NUMBER: 2003:26341 USPATFULL

Compositions for regeneration and repair of cartilage TITLE:

lesions

Atkinson, Brent, Lakewood, CO, United States INVENTOR(S):

Benedict, James J., Arvada, CO, United States

Sulzer Biologics, Inc., Austin, TX, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE

PATENT INFORMATION:

US 6511958

B1 20030128

APPLICATION INFO.:

US 2000-505209

20000216 (9)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1999-250370, filed

on 16 Feb 1999 Continuation-in-part of Ser. No. WO

1998-EP5100, filed on 12 Aug 1998

DOCUMENT TYPE:

Utility

FILE SEGMENT:

GRANTED

PRIMARY EXAMINER: LEGAL REPRESENTATIVE: Sheridan Ross P.C.

Baker, Anne-Marie

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

41 1

NUMBER OF DRAWINGS:

14 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### => d his

L3

L7

(FILE 'HOME' ENTERED AT 10:26:49 ON 28 APR 2003)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, SCISEARCH, FSTA, BIOSIS, BIOBUSINESS, WPIDS, CEABA-VTB, JAPIO, JICST-EPLUS' ENTERED AT 10:27:23 ON 28 APR 2003

3240 S OSTEOGENIC PROTEIN L1

25957 S BMP OR BONE MORPHOGENETIC PROTEIN L2

3010 S BONE MORPHOGENIC PROTEIN

1.4 2062 S L2 AND L3

302 S L1 AND L4 L5

14 S CHONDROGENIC PROTEINS 1.6

237 S CARTILAGE REPLACEMENT

1 S L7 AND L5 L8

851 S BMP-5 L9

2 S L9 AND L7 L10

762 S BMP-3 L11

2 S L11 AND L7 L12

=> s 15 and cartilage repair

14 L5 AND CARTILAGE REPAIR L13

# => d l13 ti abs ibib tot

### L13 ANSWER 1 OF 14 USPATFULL

TΤ Matrix-free osteogenic devices, implants and methods of use thereof AB Provided herein are methods for inducing bone formation in a mammal sufficient to fill a defect defining a void, wherein osteogenic protein is provided alone or dispersed in a biocompatible non-rigid, amorphous carrier having no defined surfaces. The methods and devices provide injectable formulations for filling critical size defects, as well as for accelerating the rate and enhancing the quality of bone formation in non-critical size defects.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2002:172320 USPATFULL

TITLE:

Matrix-free osteogenic devices, implants and methods of

use thereof

INVENTOR(S): Rueger, David C., Southborough, MA, UNITED STATES

Tucker, Marjorie M., Holliston, MA, UNITED STATES

PATENT ASSIGNEE(S): STRYKER CORPORATION (U.S. corporation)

APPLICATION INFO.: US 2001-887901 A1 20010622 (9)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1998-19339, filed on 5 Feb

1998, UNKNOWN

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: FISH & NEAVE, 1251 AVENUE OF THE AMERICAS, 50TH FLOOR,

NEW YORK, NY, 10020-1105

NUMBER OF CLAIMS: 37 EXEMPLARY CLAIM: 1 LINE COUNT: 2801

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 2 OF 14 USPATFULL

TI Generating cartilage in a mammal using fibroblasts transfected with a

vector encoding TGF-.beta.-1

The subject invention is related to a cell-mediated gene therapy AΒ treatment for orthopedic disease using a member belonging to the transforming growth factor..beta. (TGF-.beta.) superfamily. TGF-.beta. gene therapy as a new treatment method for degenerative arthritis is demonstrated. After transfection of TGF-.beta. cDNA expression vectors into fibroblasts (NIH 3T3-TGF-.beta.1), the cells were injected into rabbit achilles tendon and knee joints with artificially-made cartilage defects. Intratendinous injections were performed to determine the optimal concentration for in vivo expression. Partially defected cartilage model was made to simulate degenerative arthritis of the knee joint. The partial cartilage defect treated with the cell-mediated gene therapy procedure was covered by newly formed hyaline cartilage which indicates that the cells survived and stimulated matrix formation in this area. Completely denuded cartilage areas were covered by fibrous collagen.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:202190 USPATFULL

TITLE: Generating cartilage in a mammal using fibroblasts

transfected with a vector encoding TGF-.beta.-1 Noh, Moon Jong, Kyunggi-Do, Korea, Republic of

INVENTOR(S): Noh, Moon Jong, Kyunggi-Do, Korea, Republic of Kang, Kyoung Ae, Kyunggi-Do, Korea, Republic of

Lee, Kwan Hee, Seoul, Korea, Republic of

PATENT ASSIGNEE(S): TissueGene Co., Gaithersburg, MD, United States (U.S.

corporation)

DOCUMENT TYPE: Utility
FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Clark, Deborah J. R. ASSISTANT EXAMINER: Wilson, Michael C.

LEGAL REPRESENTATIVE: Squire, Sanders & Dempsey LLP., Kim, Joseph Hyosuk

NUMBER OF CLAIMS: 9 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 24 Drawing Figure(s); 9 Drawing Page(s)

LINE COUNT: 1136

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 3 OF 14 USPATFULL

Matrix-free osteogenic devices, implants and methods of use thereof Provided herein are methods for inducing bone formation in a mammal sufficient to fill a defect defining a void, wherein osteogenic protein is provided alone or dispersed in a biocompatible non-rigid, amorphous carrier having no defined surfaces. The methods and devices provide injectable formulations for filling critical size defects, as well as for accelerating the rate and enhancing the quality of bone formation in non-critical size defects.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:142331 USPATFULL

TITLE: Matrix-free osteogenic devices, implants and methods of

use thereof

INVENTOR(S): Rueger, David C., Southborough, MA, United States

Tucker, Marjorie M., Holliston, MA, United States

PATENT ASSIGNEE(S): Stryker Corporation, Kalamazoo, MI, United States (U.S.

corporation)

FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Russel, Jeffrey E.

LEGAL REPRESENTATIVE: Fish & Neave, Haley, Jr., James F., Mangasarian, Karen

NUMBER OF CLAIMS: 25 EXEMPLARY CLAIM: 1 LINE COUNT: 2501

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 4 OF 14 USPATFULL

TI OSTEOGENIC DEVICES AND METHODS OF USE THEREOF FOR REPAIR OF ENDOCHONDRAL BONE, OSTEOCHONDRAL AND CHONDRAL DEFECTS

Disclosed herein are improved osteogenic devices and methods of use thereof for repair of bone and cartilage defects. The devices and methods promote accelerated formation of repair tissue with enhanced stability using less osteogenic protein than devices in the art. Defects susceptible to repair with the instant invention include, but are not limited to: critical size defects, non-critical size defects, non-union fractures, fractures, osteochondral defects, subchondral defects, and defects resulting from degenerative diseases such as osteochondritis dessicans.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:139603 USPATFULL

TITLE: OSTEOGENIC DEVICES AND METHODS OF USE THEREOF FOR

REPAIR OF ENDOCHONDRAL BONE, OSTEOCHONDRAL AND CHONDRAL

DEFECTS

INVENTOR(S): RUEGER, DAVID C., SOUTHBOROUGH, MA, United States

TUCKER, MARJORIE A., HOLLISTON, MA, United States CHANG, AN-CHENG, WESTBOROUGH, MA, United States

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: PATENT ADMINISTATOR, TESTA HURWITZ & THIBEAULT, LLP,

HIGH STREET TOWER, 125 HIGH STREET, BOSTON, MA, 02110

NUMBER OF CLAIMS: 49

EXEMPLARY CLAIM:

1

NUMBER OF DRAWINGS:

2 Drawing Page(s)

LINE COUNT:

5269

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 5 OF 14 USPATFULL

TI IMPROVED OSTEOGENIC DEVICES AND METHODS OF USE THEREOF FOR REPAIR OF ENDOCHONDRAL BONE AND OSTEOCHONDRAL DEFECTS

Disclosed herein are improved osteogenic devices and methods of use thereof for repair of bone and cartilage defects. The devices and methods promote accelerated formation of repair tissue with enhanced stability using less osteogenic protein than devices in the art. Defects susceptible to repair with the instant invention include, but are not limited to: critical size defects, non-critical size defects, non-union fractures, fractures, osteochondral defects, subchondral defects, and defects resulting from degenerative diseases such as osteochondritis dessicans.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2001:134213 USPATFULL

TITLE:

IMPROVED OSTEOGENIC DEVICES AND METHODS OF USE THEREOF

FOR REPAIR OF ENDOCHONDRAL BONE AND OSTEOCHONDRAL

**DEFECTS** 

INVENTOR(S):

RUEGER, DAVID C, SOUTHBOROUGH, MA, United States TUCKER, MARJORIE A, HOLLISTON, MA, United States

	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 2001014662	A1	20010816	
APPLICATION INFO.:	US 1997-822186	A1	19970320	(8)
DOCUMENT TYPE:	Utility			

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

JAMES F. HALEY, FISH & NEAVE, 1251 AVENUE OF THE

AMERICAS, NEW YORK, NY, 100201104

NUMBER OF CLAIMS: 34
EXEMPLARY CLAIM: 1

EXEMPLARY CLAIM: 1
NUMBER OF DRAWINGS: 2

2 Drawing Page(s)

LINE COUNT:

4425

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

## L13 ANSWER 6 OF 14 USPATFULL

TI Methods and compositions for the treatment and repair of defects or lesions in cartilage or bone using functional barrier

Methods and compositions are provided for the treatment and repair of AB defects in the cartilage or bone of humans and other animals as in full-thickness defects in joints. To induce cartilage formation, a defect in cartilage is filled with a matrix having pores sufficiently large to allow cartilage repair cells to populate the matrix. The matrix contains an anti-angiogenic agent that serves as a functional barrier to prevent vascularization and bone growth into the cartilage area. The matrix filling the defect in cartilage may also contain a proliferation agent and a chemotactic agent, and a transforming factor in an appropriate delivery system. A functional barrier between the bone and cartilage areas of a full-thickness defect may also be created by heat-treating the areas of bleeding to form a transient tissue barrier which prevents blood vessels and associated cells from penetrating from the bone area into the cartilage area. If desired, the bone portion of the full-thickness defect may be filled with a matrix having pores large enough to allow cells to populate the matrix and to form blood vessels. The matrix filling the bone defect may contain an angiogenic factor and an osteogenic factor in an appropriate delivery system. Methods and compositions are also provided for assisted bone and connective tissue regeneration for dental and other applications.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

1998:162021 USPATFULL

TITLE:

Methods and compositions for the treatment and repair

of defects or lesions in cartilage or bone using

functional barrier

INVENTOR(S):

Hunziker, Ernst B., Riedholz, Switzerland

PATENT ASSIGNEE(S):

Shaw, Robert Francis, Sausalito, CA, United States

(U.S. individual)

NUMBER KIND DATE

PATENT INFORMATION:

19981229

APPLICATION INFO.:

US 5853746 US 1996-672618

19960628 (8)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1995-524034, filed on 6 Sep 1995, now abandoned which is a continuation of Ser. No. US 1994-338126, filed on 1 Nov 1994, now

abandoned which is a continuation of Ser. No. US 1992-979904, filed on 23 Nov 1992, now patented, Pat. No. US 5368858 which is a division of Ser. No. US 1991-648274, filed on 31 Jan 1991, now patented, Pat.

No. US 5206023

DOCUMENT TYPE:

Utility

FILE SEGMENT:

Granted

PRIMARY EXAMINER:

Azpuru, Carlos A.

LEGAL REPRESENTATIVE: Fish & Neave, Massaro, Jane A., Rosen, Mark J.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

30

LINE COUNT:

1673

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

## L13 ANSWER 7 OF 14 USPATFULL

ΤI Methods and compositions for the treatment and repair of defects or lesions in cartilage or bone

Methods and compositions are provided for the treatment and repair of AB defects in the cartilage or bone of humans and other animals as in full-thickness defects in joints. The defect in bone is filled with a matrix having pores large enough to allow cells to populate the matrix and to form blood vessels. The matrix filling the bone defect contains an angiogenic factor and also contains an osteogenic factor in an appropriate delivery system. To induce cartilage formation, a defect in cartilage is filled with a matrix having pores sufficiently large to allow cartilage repair cells to populate the matrix.

The matrix filling the defect in cartilage contains a proliferation agent and also contains a transforming factor in an appropriate delivery system. The matrix may also contain a chemotactic agent to attract cartilage repair cells. In a full-thickness defect,

the defect sites in bone and cartilage are separated from each other by a membrane, which is sealed to the cartilage-bone-junction and which prevents blood vessels and associated cells from penetrating from one site to the other.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

93:104945 USPATFULL

TITLE:

Methods and compositions for the treatment and repair

of defects or lesions in cartilage or bone

INVENTOR(S):

Hunziker, Ernst B., Riedholz, Switzerland

PATENT ASSIGNEE(S):

Shaw, Robert Francis, San Francisco, CA, United States

(U.S. individual)

NUMBER KIND DATE PATENT INFORMATION: US 5270300 19931214 APPLICATION INFO.: US 1991-756164 19910906 (7) DOCUMENT TYPE:

Utility Granted

FILE SEGMENT: PRIMARY EXAMINER:

Griffin, Ronald W.

LEGAL REPRESENTATIVE:

Mullowney, Edward F., Massaro, Jane A.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

26 1,10

LINE COUNT:

1089

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 8 OF 14 DGENE (C) 2003 THOMSON DERWENT L13

Implant for inducing local bone or cartilage formation, comprising TТ

osteogenic proteins, non-synthetic polymeric matrixes and binding agents

AAB82698 Protein DGENE AN

AB The present sequence is that of osteogenic protein generic sequence 8, which accommodates the homologies shared between osteogenic proteins OP-1, OP-2, OP-3, CBMP-2A, CBMP-2B, bone morphogenic protein (BMP)-3, Drosophila

protein 60A, DPP, Vgl, BMP-5, BMP-6, Vgr-1 and GDF-1. The generic sequence includes a 7 cysteine domain, providing an appropriate cysteine skeleton for the formation of inter- or intramolecular disulfide bonds, and also includes certain critical amino acids likely to influence the tertiary structure of folded proteins. Provision of an 8th cysteine residue at position 41 encompasses the morphogenically active sequences of OP-2 and OP-3. Proteins based on the present sequence can be used in novel osteogenic devices of the invention. The invention is based on the discovery that admixing osteogenic protein and a non-synthetic, non-polymeric matrix such as collagen or beta-tricalcium phosphate with a binding agent yields an improved osteogenic device with enhanced bone and cartilage repair capabilities. The osteogenic devices

promote accelerated formation of repair tissue with enhanced stability using less osteogenic protein than previous devices. Defects susceptible to repair include critical size defects, non-critical size defects, non-union fractures, fractures, osteochondral defects and

subchondral defects (claimed), and defects resulting from degenerative

diseases such as osteochondritis desiccans.

ACCESSION NUMBER: AAB82698 Protein **DGENE** 

TITLE:

Implant for inducing local bone or cartilage formation, comprising osteogenic proteins, non-synthetic polymeric

matrixes and binding agents -

Rueger D C; Tucker M A; Chang A INVENTOR:

PATENT ASSIGNEE: (RUEG-I) RUEGER D C.

(TUCK-I) TUCKER M A. (CHAN-I) CHANG A.

PATENT INFO: US 2001016646 A1 20010823 59p

APPLICATION INFO: US 1998-45331 19980320 PRIORITY INFO: US 1998-45331 19980320

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 2001-513983 [56]

DESCRIPTION: Osteogenic protein generic sequence 8.

- ANSWER 9 OF 14 DGENE (C) 2003 THOMSON DERWENT L13
- ΤI Implant for inducing local bone or cartilage formation, comprising osteogenic proteins, non-synthetic polymeric matrixes and binding agents
- ΑN AAB82697 Protein DGENE
- AB The present sequence is that of osteogenic protein generic sequence 7, which accommodates the homologies shared between osteogenic proteins OP-1, OP-2, OP-3, CBMP-2A, CBMP-2B, bone morphogenic protein (BMP)-3, Drosophila protein 60A, DPP, Vgl, BMP-5, BMP-6, Vgr-1 and GDF-1. The generic sequence includes a 6 cysteine domain, providing an

appropriate cysteine skeleton for the formation of inter- or intramolecular disulfide bonds, and also includes certain critical amino acids likely to influence the tertiary structure of folded proteins. Provision of a 7th cysteine residue at position 36 encompasses the morphogenically active sequences of OP-2 and OP-3. Proteins based on the present sequence can be used in novel osteogenic devices of the invention. The invention is based on the discovery that admixing osteogenic protein and a non-synthetic, non-polymeric matrix such as collagen or beta-tricalcium phosphate with a binding agent yields an improved osteogenic device with enhanced bone and cartilage repair capabilities. The osteogenic devices promote accelerated formation of repair tissue with enhanced stability using less osteogenic protein than previous devices. Defects susceptible to repair include critical size defects, non-critical size defects, non-union fractures, fractures, osteochondral defects and subchondral defects (claimed), and defects resulting from degenerative diseases such as osteochondritis desiccans.

ACCESSION NUMBER: AAB82697 Protein

Implant for inducing local bone or cartilage formation, TITLE:

comprising osteogenic proteins, non-synthetic polymeric

matrixes and binding agents -

Rueger D C; Tucker M A; Chang A INVENTOR:

PATENT ASSIGNEE: (RUEG-I) RUEGER D C.

(TUCK-I) TUCKER M A. (CHAN-I) CHANG A.

US 2001016646 A1 20010823 59p PATENT INFO:

APPLICATION INFO: US 1998-45331 19980320 PRIORITY INFO: US 1998-45331 19980320

DOCUMENT TYPE: Patent English

OTHER SOURCE: 2001-513983 [56]

DESCRIPTION: Osteogenic protein generic sequence 7.

ANSWER 10 OF 14 DGENE (C) 2003 THOMSON DERWENT L13

Implant for inducing local bone or cartilage formation, comprising TIosteogenic proteins, non-synthetic polymeric matrixes and binding agents

AAB82695 Protein DGENE AN

The present sequence is that of human osteogenic AB protein OP-1. The invention is based on the discovery that admixing osteogenic protein and a non-synthetic, non-polymeric matrix such as collagen or beta-tricalcium phosphate with a

binding agent yields an improved osteogenic device with enhanced bone and cartilage repair capabilities. The osteogenic

protein may be OP-1, OP-2, bone morphogenic

protein (BMP)-2, BMP-4, BMP-5, BMP-6, BMP-9, BMP-10, BMP-11,

BMP-12, BMP-15, BMP-16, DPP, Vgl, Vgr, 60A

protein, GDF1, GDF3, GDF5, GDF6, GDF7, GDF8, GDF9, GDF10, GDF11, or their variants, and is especially OP-1. The osteogenic devices promote accelerated formation of repair tissue with enhanced stability using less

osteogenic protein than previous devices. Defects

susceptible to repair include critical size defects, non-critical size defects, non-union fractures, fractures, osteochondral defects and subchondral defects (claimed), and defects resulting from degenerative diseases such as osteochondritis desiccans.

ACCESSION NUMBER: AAB82695 Protein DGENE

TITLE: Implant for inducing local bone or cartilage formation,

comprising osteogenic proteins, non-synthetic polymeric

matrixes and binding agents -

INVENTOR: Rueger D C; Tucker M A; Chang A PATENT ASSIGNEE: (RUEG-I) RUEGER D C.

(TUCK-I) TUCKER M A. (CHAN-I) CHANG A.

PATENT INFO: US 2001016646 A1 20010823 59p

APPLICATION INFO: US 1998-45331 19980320 PRIORITY INFO: US 1998-45331

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 2001-513983 [56] CROSS REFERENCES: N-PSDB: AAH26404

DESCRIPTION: Human osteogenic protein OP-1.

ANSWER 11 OF 14 DGENE (C) 2003 THOMSON DERWENT L13

Implant for inducing local bone or cartilage formation, comprising TΤ osteogenic proteins, non-synthetic polymeric matrixes and binding agents

AN AAH26404 cDNA **DGENE** 

The present sequence is that of cDNA encoding human osteogenic AΒ protein OP-1 (see AAB82695). The invention is based on the discovery that admixing osteogenic protein and a non-synthetic, non-polymeric matrix such as collagen or beta-tricalcium phosphate with a binding agent yields an improved osteogenic device with enhanced bone and cartilage repair capabilities. The osteogenic protein may be OP-1, OP-2, bone

morphogenic protein (BMP)-2, BMP-4,

BMP-5, BMP-6, BMP-9, BMP-10, BMP-11, BMP-12, BMP-15, BMP-16,

DPP, Vgl, Vgr, 60A protein, GDF1, GDF3, GDF5, GDF6, GDF7, GDF8, GDF9, GDF10, GDF11, or their variants, and is especially OP-1. The osteogenic devices promote accelerated formation of repair tissue with enhanced stability using less osteogenic protein than previous devices. Defects susceptible to repair include critical size defects, non-critical size defects, non-union fractures, fractures, osteochondral defects and subchondral defects (claimed), and defects resulting from degenerative diseases such as osteochondritis desiccans.

ACCESSION NUMBER: AAH26404 cDNA DGENE

TITLE: Implant for inducing local bone or cartilage formation,

comprising osteogenic proteins, non-synthetic polymeric

59p

matrixes and binding agents -

INVENTOR: Rueger D C; Tucker M A; Chang A

PATENT ASSIGNEE: (RUEG-I) RUEGER D C.

(TUCK-I) TUCKER M A.

(CHAN-I) CHANG A. INFO: US 2001016646 A1 20010823 PATENT INFO:

APPLICATION INFO: US 1998-45331 19980320 PRIORITY INFO: US 1998-45331
DOCUMENT TYPE: Datent 19980320

DOCUMENT TYPE: Patent

LANGUAGE: English
OTHER SOURCE: 2001-513983 [56] CROSS REFERENCES: P-PSDB: AAB82695

DESCRIPTION: Human osteogenic protein OP-1 cDNA.

L13 ANSWER 12 OF 14 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.

ТΤ Enhanced matrix synthesis and in vitro formation of cartilage-like tissue by genetically modified chondrocytes expressing BMP-7.

Bone morphogenic protein-7 (BMP-7) AB

supports ectopic cartilage and bone formation, is expressed in normal articular cartilage, and increases matrix synthesis in chondrocytes. Based on this knowledge, we hypothesized that an adenovirus (Ad) vector encoding human BMP-7 could be used to modify chondrocytes genetically to improve their capacity for cartilage repair. An adenovirus vector encoding BMP-7 (AdBMP-7) was constructed and its bioactivity confirmed by ectopic bone formation assay. AdBMP-7 modification of bovine chondrocytes induced expression of BMP-7 mRNA and bioactive protein, resulting in an increase in incorporation of (35)SO(-)(4) into proteoglycan, (3)H-proline uptake into protein, and the expression of the cartilage-specific matrix genes, aggrecan and type II

collagen. An in vitro model of chondrocyte transplantation was used to demonstrate the feasibility of using genetically modified chondrocytes to enhance formation of cartilage-like tissue. When transplanted onto cartilage explants and maintained in vitro for 3 weeks, chondrocytes modified with AdBMP-7 formed 1.9-fold thicker tissue than chondrocytes modified with a control vector (P < 0.001). This tissue was positive for type II collagen and proteoglycan but negative for type X collagen and demonstrated a cartilage-like morphology. These observations suggest that Ad-mediated transfer of BMP-7 gene to chondrocytes enhances the chondrocyte-specific matrix synthesis and their capacity to form cartilage-like tissue, thus representing a strategy that may improve cell-based cartilage repair. . COPYRGT. 2001.

Orthopaedic Research Society. Published by Elsevier Science Ltd. All rights reserved.

2001305182 EMBASE ACCESSION NUMBER:

Enhanced matrix synthesis and in vitro formation of TITLE:

cartilage-like tissue by genetically modified chondrocytes

expressing BMP-7.

Hidaka C.; Quitoriano M.; Warren R.F.; Crystal R.G. AUTHOR: C. Hidaka, Institute of Genetic Medicine, Weill Medical CORPORATE SOURCE:

Coll. of Cornell Univ., New York, NY 10021, United States.

geneticmedicine@mail.med.cornell.edu

SOURCE: Journal of Orthopaedic Research, (2001) 19/5 (751-758).

Refs: 40

ISSN: 0736-0266 CODEN: JOREDR

S 0736-0266(01)00019-5 PUBLISHER IDENT.:

COUNTRY:

United Kingdom

DOCUMENT TYPE:

Journal; Article 004 Microbiology

FILE SEGMENT: 022 Human Genetics

> Clinical Biochemistry 029 Orthopedic Surgery 033

LANGUAGE:

SUMMARY LANGUAGE: English

L13 ANSWER 13 OF 14 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.

Expression of human bone morphogenic protein

English

7 in primary rabbit periosteal cells: Potential utility in gene therapy for osteochondral repair.

A commonly encountered problem in orthopedics is bone and cartilage tissue AΒ injury which heals incompletely or without full structural integrity. This necessitates development of improved methods for treatment of injuries which are not amenable to treatment using current therapies. An already large and growing number of growth factors which play significant roles in bone remodeling and repair have been identified in the past few years. It is well established that bone morphogenic proteins induce the production of new bone and cartilage. An efficient method of delivery of these growth factors by conventional pharmacological means has yet to be elucidated. We wished to evaluate the use of retroviral vector-mediated gene transfer to deliver genes of therapeutic relevance for bone and cartilage repair. To determine the feasibility of using amphotropically packaged retroviral vectors to transduce primary rabbit mesenchymal stem cells of periosteal origin, primary periosteal cells were isolated from New Zealand white rabbits, transduced in vitro with a retroviral vector bearing both the nuclear localized lacZ marker gene and the neo(r) gene, and selected in G418. We used a convenient model for analysis of in vivo stability of these cells which were seeded on to polymer scaffold grafts and implanted into rabbit femoral osteochondral defects. The nuclear localized .beta.-galactosidase protein was expressed in essentially 100% of selected cells in vitro and was observed in the experimental explants from animals after both 4 and 8 weeks in vivo, while cells transduced with a retroviral vector bearing only the neo(r) gene in negative control explants showed no blue staining. We extended our study by delivering a gene of therapeutic relevance, human bone morphogenic

protein 7 (hBMP-7), to primary periosteal cells via retroviral vector. The hBMP-7 gene was cloned from human kidney 293 cell total RNA by RT-PCR into a retroviral vector under control of the CMV enhancer/promoter. Hydroxyapatite secretion, presumably caused by overexpression of hBMP-7, was observed on the surface of the transduced and selected periosteal cells, however, this level of expression was toxic to both PA317 producer and primary periosteal cells. Subsequently, the strong CMV enhancer/promoter driving the hBMP-7 gene was replaced in the retroviral vector by a weaker enhancer/promoter from the rat .beta.-actin gene. Non-toxic levels of expression of hBMP-7 were confirmed at both the RNA and protein levels in PA317 producer and primary periosteal cell lines and cell supernatants. This work demonstrates the feasibility of using a gene therapy approach in attempts to promote bone and cartilage tissue repair using gene-modified periosteal cells on grafts.

ACCESSION NUMBER: 1998294320 EMBASE

TITLE: Expression of human bone morphogenic

protein 7 in primary rabbit periosteal cells:

Potential utility in gene therapy for osteochondral repair. Mason J.M.; Grande D.A.; Barcia M.; Grant R.; Pergolizzi

AUTHOR: Mason J.M.; Grande D R.G.; Breitbart A.S.

CORPORATE SOURCE: J.M. Mason, Viral Vector Laboratory, Department of

Research, North Shore University Hospital, 350 Community

Drive, Manhasset, NY 11030, United States

SOURCE: Gene Therapy, (1998) 5/8 (1098-1104).

Refs: 27

ISSN: 0969-7128 CODEN: GETHEC

COUNTRY: United Kingdom DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 033 Orthopedic Surgery

LANGUAGE: English SUMMARY LANGUAGE: English

AB

L13 ANSWER 14 OF 14 SCISEARCH COPYRIGHT 2003 THOMSON ISI

TI Expression of human bone morphogenic protein

7 in primary rabbit periosteal cells: potential utility in gene therapy for osteochondral repair

A commonly encountered problem in orthopedics is bone and cartilage tissue injury which heals incompletely or without full structural integrity. This necessitates development of improved methods for treatment of injuries which are not amenable to treatment using current therapies. An already large and growing number of growth factors which play significant roles in bone remodeling and repair have been identified in the past few years. It is well established that bone morphogenic proteins induce the production of new bone and cartilage. An efficient method of delivery of these growth factors by conventional pharmacological means has yet to be elucidated We wished to evaluate the use of retroviral vector-mediated gene transfer to deliver genes of therapeutic relevance for bone and cartilage repair. To determine the feasibility of using amphotropically packaged retroviral vectors to transduce primary rabbit mesenchymal stem cells of periosteal origin, primary periosteal cells were isolated from New Zealand white rabbits, transduced in vitro with a retroviral vector bearing both the nuclear localized lacZ marker gene and the neo(r) gene, and selected in G418. We used a convenient model for analysis of in vivo stability of these cells which were seeded on to polymer scaffold grafts and implanted into rabbit femoral osteochondral defects. The nuclear localized beta-galactosidase protein was expressed in essentially 100% of selected cells in vitro and was observed in the experimental explants from animals after both 4 and 8 weeks in vivo, while cells transduced with a retroviral vector bearing only the neo(r) gene in negative control explants showed no blue staining. We extended our study by delivering a gene of therapeutic relevance, human bone morphogenic protein 7 (hBMP-7), to primary periosteal cells via retroviral vector. The hBMP-7 gene was cloned

from human kidney 293 cell total RNA by RT-PCR into a retroviral vector

under control of the CMV enhancer/promoter. Hydroxyapatite secretion, presumably caused by overexpression of hBMP-7, was observed on the surface of the transduced and selected periosteal cells, however, this level of expression was toxic to both PA317 producer and primary periosteal cells. Subsequently, the strong CMV enhancer/promoter driving the hBMP-7 gene was replaced in the retroviral vector by a weaker enhancer/promoter from the rat beta-actin gene. Nontoxic levels of expression of hBMP-7 were confirmed at both the RNA and protein levels in PA317 producer and primary periosteal cell lines and cell supernatants. This work demonstrates the feasibility of using a gene therapy approach in attempts to promote bone and cartilage tissue repair using gene-modified periosteal cells on grafts.

ACCESSION NUMBER:

1998:612330 SCISEARCH

THE GENUINE ARTICLE: 107RU

TITLE:

Expression of human bone morphogenic

protein 7 in primary rabbit periosteal cells:

potential utility in gene therapy for osteochondral repair

AUTHOR: Mason J M (Reprint); Grande D A; Barcia M; Grant R;

Pergolizzi R G; Breitbart A S

CORPORATE SOURCE:

N SHORE UNIV HOSP, NYU, SCH MED, DEPT RES, VIRAL VECTOR LAB, 350 COMMUNITY DR, MANHASSET, NY 11030 (Reprint); N SHORE UNIV HOSP, NYU, SCH MED, DEPT SURG, DIV ORTHOPED, MANHASSET, NY 11030; N SHORE UNIV HOSP, NYU, SCH MED, DEPT SURG, DIV PLAST & RECONSTRUCT SURG, MANHASSET, NY 11030

COUNTRY OF AUTHOR:

USA

SOURCE:

GENE THERAPY, (AUG 1998) Vol. 5, No. 8, pp. 1098-1104. Publisher: STOCKTON PRESS, HOUNDMILLS, BASINGSTOKE RG21

6XS, HAMPSHIRE, ENGLAND.

ISSN: 0969-7128.

DOCUMENT TYPE: '

Article; Journal

FILE SEGMENT:

LIFE

LANGUAGE:

English

REFERENCE COUNT:

27
\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

=> s articular

L14 127781 ARTICULAR

=> s nonarticular cartilage

L15 30 NONARTICULAR CARTILAGE

=> d 115 and defect locus

'AND' IS NOT A VALID FORMAT

'DEFECT' IS NOT A VALID FORMAT

'LOCUS' IS NOT A VALID FORMAT

In a multifile environment, a format can only be used if it is valid in at least one of the files. Refer to file specific help messages or the STNGUIDE file for information on formats available in individual files.

REENTER DISPLAY FORMAT FOR ALL FILES (FILEDEFAULT): end

=> s l15 and defect locus

L16 9 L15 AND DEFECT LOCUS

=> d l16 ti abs ibib tot

L16 ANSWER 1 OF 9 USPATFULL

TI Repair of larynx, trachea, and other fibrocartilaginous tissues

AB Provided herein are methods and devices for inducing the formation of functional replacement nonarticular cartilage tissues and ligament tissues. These methods and devices involve the use of osteogenic proteins, and are useful in repairing defects in the larynx, trachea, interarticular menisci, intervertebral discs, ear,

nose, ribs and other fibrocartilaginous tissues in a mammal.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2001:165613 USPATFULL ACCESSION NUMBER:

Repair of larynx, trachea, and other fibrocartilaginous TITLE:

Vukicevic, Slobodan, Zagreb, Croatia INVENTOR(S):

Katic, Vladimir, Zagreb, Croatia

Sampath, Kuber T., Holliston, MA, United States

Creative BioMolecules, Inc. (non-U.S. corporation) PATENT ASSIGNEE(S):

> NUMBER KIND DATE \_\_\_\_\_\_

US 2001024823 A1 20010927 US 2001-828607 A1 20010406 (9) PATENT INFORMATION: PATENT INFORMATION:
APPLICATION INFO.:

RELATED APPLN. INFO.: Continuation of Ser. No. WO 1999-US17222, filed on 30

Jul 1999, UNKNOWN

NUMBER DATE -----

PRIORITY INFORMATION: US 1998-103161P 19981006 (60)

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: FISH & NEAVE, 1251 AVENUE OF THE AMERICAS, 50TH FLOOR,

NEW YORK, NY, 10020-1105

NUMBER OF CLAIMS: 56 EXEMPLARY CLAIM: 1 LINE COUNT: 1859

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 2 OF 9 DGENE (C) 2003 THOMSON DERWENT L16

ΤI Novel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AN AAY92442 Protein DGENE

The specification concerns a novel method for repairing a defect in a AB non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of

functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92442 Protein DGENE

TITLE: Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable

carrier

INVENTOR: Vukicevic S; Katic V; Sampath K T

PATENT ASSIGNEE: (STYC)STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413 APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006
DOCUMENT TYPE: Patent
LANGUAGE: English
OTHER SOURCE: 2000-317644 [27]

CROSS REFERENCES: N-PSDB: AAA09361

DESCRIPTION: Human osteogenic protein 1 (OP-1).

ANSWER 3 OF 9 DGENE (C) 2003 THOMSON DERWENT L16

TI Novel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AAY92441 protein DGENE AN

Generic Sequence 10 contains generic sequence 9 and an N-terminal AB extension. Generic sequence 9 is a composite amino acid sequence of the following proteins: human OP-1 to -3, human BMP-2 to -6, -9 to -11, Drosophila 60A, Xenopus Vq-1, sea urchin UNIVIN, human CDMP-1 to -3, human and mouse GDF-1, chicken DORSALIN, DPP, Drosophila Screw, mouse NODAL, mouse GDF-8 to -11, human GDF-8, -11, human BMP-15 and rat BMP3b. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92441 protein **DGENE** 

TITLE: Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable

carrier

Vukicevic S; Katic V; Sampath K T INVENTOR:

PATENT ASSIGNEE: (STYC) STRYKER CORP.

WO 2000020021 A1 20000413 65p PATENT INFO:

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent LANGUAGE: English

2000-317644 [27]

LANGUAGE.
OTHER SOURCE: DESCRIPTION: Generic sequence 10, derived from osteogenic protein family

ANSWER 4 OF 9 DGENE (C) 2003 THOMSON DERWENT L16

Novel methods for repairing a defect in mammalian nonarticular ΤI cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

ANAAY92440 protein DGENE

Generic Sequence 9 is a composite amino acid sequence of the following AB proteins: human OP-1 to -3, human BMP-2 to -6, -9 to -11, Drosophila 60A, Xenopus Vg-1, sea urchin UNIVIN, human CDMP-1 to -3, human and mouse GDF-1, chicken DORSALIN, DPP, Drosophila Screw, mouse NODAL, mouse GDF-8 to -11, human GDF-8, -11, human BMP-15 and rat BMP3b. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92440 protein DGENE

Novel methods for repairing a defect in mammalian TITLE:

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable

carrier

Vukicevic S; Katic V; Sampath K T INVENTOR:

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413 65p

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent English LANGUAGE:

OTHER SOURCE: 2000-317644 [27]

Generic sequence 9, derived from osteogenic protein family DESCRIPTION:

members.

ANSWER 5 OF 9 DGENE (C) 2003 THOMSON DERWENT L16

Novel methods for repairing a defect in mammalian nonarticular TΤ cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

**DGENE** ANAAY92439 protein

Generic Sequence 8 contains generic sequence 7 (AAY92438), which AB accomodates the homologies shared among osteogenic protein family members, including OP-1, OP-2, OP-3, BMP-2 to -6, 60A, DPP, Vg-1, Vgr-1 and GDF, as well as an N-terminal addition of 5 residues. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs,

**DGENE** ACCESSION NUMBER: AAY92439 protein

Novel methods for repairing a defect in mammalian TITLE:

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable

carrier

Vukicevic S; Katic V; Sampath K T INVENTOR:

PATENT ASSIGNEE: (STYC) STRYKER CORP.

and interarticular menisci.

PATENT INFO: WO 2000020021 A1 20000413 65p

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2000-317644 [27]

DESCRIPTION: Generic sequence 8, derived from osteogenic protein family

members.

ANSWER 6 OF 9 DGENE (C) 2003 THOMSON DERWENT L16

Novel methods for repairing a defect in mammalian nonarticular TТ cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AAY92438 protein ΆN DGENE

AB Generic Sequence 7 accomodates the homologies shared among osteogenic protein family members, including OP-1, OP-2, OP-3, BMP-2 to -6, 60A, DPP, Vg-1, Vgr-1 and GDF. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the

glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92438 protein DGENE

TITLE: Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable

carrier

INVENTOR: Vukicevic S; Katic V; Sampath K T

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413 65p

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 2000-317644 [27]

DESCRIPTION: Generic sequence 7, derived from osteogenic protein family

members.

L16 ANSWER 7 OF 9 DGENE (C) 2003 THOMSON DERWENT

Novel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AN AAY92437 protein DGENE

OPX defines the seven-cysteine skeleton of several OP-1 and OP-2 AΒ variants. Each Xaa is chosen from the residues occuring at the corresponding position in the C-terminal sequence of mouse or human OP-1 or OP-2. The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAY92437 protein DGENE

TITLE: Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable

carrier

INVENTOR: Vukicevic S; Katic V; Sampath K T

PATENT ASSIGNEE: (STYC) STRYKER CORP.

PATENT INFO: WO 2000020021 A1 20000413 65p

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 2000-317644 [27]

DESCRIPTION: Generic OPX, seven-cysteine skeleton.

L16 ANSWER 8 OF 9 DGENE (C) 2003 THOMSON DERWENT

Novel methods for repairing a defect in mammalian nonarticular cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier

AN AAA09361 cDNA DGENE

AB The specification concerns a novel method for repairing a defect in a non-articular cartilage tissue or a ligament of a mammal, which comprises providing an osteogenic protein in a biocompatible, bioresorbable carrier

to the defect locus to induce the formation of functional replacement cartilage. The methods and implants, promote chondrogenesis and are useful for repairing or correcting a defect in a non-articular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, oedema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci.

ACCESSION NUMBER: AAA09361 cDNA

Novel methods for repairing a defect in mammalian TITLE:

nonarticular cartilage tissue or ligaments

using an osteogenic protein in a biocompatible, bioresorbable

carrier

Vukicevic S; Katic V; Sampath K T INVENTOR:

PATENT ASSIGNEE: (STYC) STRYKER CORP.

WO 2000020021 A1 20000413 65p PATENT INFO:

APPLICATION INFO: WO 1999-US17222 19990730 PRIORITY INFO: US 1998-103161 19981006

DOCUMENT TYPE: Patent

LANGUAGE: English
OTHER SOURCE: 2000-317644 [27] CROSS REFERENCES: P-PSDB: AAY92442

DESCRIPTION: Human osteogenic protein 1 (OP-1) coding sequence.

L16 ANSWER 9 OF 9 WPIDS (C) 2003 THOMSON DERWENT

Novel methods for repairing a defect in mammalian nonarticular TТ cartilage tissue or ligaments using an osteogenic protein in a biocompatible, bioresorbable carrier.

AN 2000-317644 [27] WPIDS

2000-317706 [27] CR

AB WO 200020021 A UPAB: 20020910

> NOVELTY - Repairing a defect in a nonarticular cartilage tissue or a ligament of a mammal, comprising providing an osteogenic protein in a biocompatible, bioresorbable carrier to the defect locus, inducing the formation of functional replacement cartilage, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) an implantable device for repairing a defect in a nonarticular cartilage tissue comprising an osteogenic protein disposed in a devitalized cartilage, a collagen carrier, or a carboxymethylcellulose carrier; and
- (2) promoting chondrogenesis at a defect locus in a mammal comprising providing an osteogenic protein in a devitalized cartilage carrier that is configured to fit into the defect locus.

ACTIVITY - Osteogenic; chondrogenic.

MECHANISM OF ACTION - Osteopathic stimulating implant; transplantation.

USE - The methods and implants are useful for repairing or correcting a defect in a nonarticular cartilage tissue or a ligament of a mammal, e.g. cleft larynx, edema of the glottis, ulceration of the larynx caused by syphilis, tuberculosis or malignancy, defects resulting from mechanical trauma to the larynx or trachea (including tracheotomy and laryngotomy), laryngeal cancer, and defects of the ear, nose, ribs, invertebral discs, and interarticular menisci. Dwg.0/0

ACCESSION NUMBER: 2000-317644 [27] WPIDS

2000-317706 [27] CROSS REFERENCE: DOC. NO. CPI: C2000-096081

TITLE: Novel methods for repairing a defect in mammalian

nonarticular cartilage tissue or

ligaments using an osteogenic protein in a biocompatible,

bioresorbable carrier.

DERWENT CLASS:

A96 B04 D22

INVENTOR(S): KATIC, V; SAMPATH, K T; VUKICEVIC, S
PATENT ASSIGNEE(S): (STYC) STRYKER CORP; (CREA-N) CREATIVE BIOMOLECULES INC

COUNTRY COUNT:

23

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2000020021 A1 20000413 (200027)\* EN 64

RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE

W: AU CA JP US

AU 9952417 A 20000426 (200036) EP 1117422 A1 20010725 (200143) EN

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US 2001024823 A1 20010927 (200159)

70 JP 2002526167 W 20020820 (200258)

## APPLICATION DETAILS:

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WO 2000020021 A1 AU 9952417 A	WO 1999-US17222 AU 1999-52417	19990730 19990730
AU 9952417 A EP 1117422 A1	EP 1999-937624 WO 1999-US17222	19990730 19990730
US 2001024823 A1 Provisional	WO 1999-US17222 US 1998-103161P WO 1999-US17222	19990730 19981006 19990730
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